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PREFACE

The work described in this report was authorized under Project IM162622A554, Chemical Munitions and Chemical Combat Support; Technical Area 4-5, Smoke Toxicology. The work was started in July 1978 and completed in May 1980. The experimental data are contained in notebook 9839.

In conducting the research described in this report, the investigators adhered to the "Guide for the Care and Use of Laboratory Animals" as promulgated by the Committee on Revision of the Guide for Laboratory Animals Facilities and Care of the Institute of Laboratory Animal Resources, National Research Council.

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This report has not been approved for release to the public.

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CONTENTS

	<i>Page</i>
1 INTRODUCTION	7
2 EXPERIMENTAL PROCEDURES	7
2.1 Chamber Operation and Sample Collection	7
2.2 Animals Used	8
2.3 Animal Exposures and Observations	8
2.4 Blood Studies	8
2.5 Pathology	8
2.6 Physiological and Behavioral Studies	9
3 RESULTS	9
3.1 Exposure Conditions	9
3.2 Toxic Signs	9
3.3 Body-Organ Weights	9
3.4 Hematology and Blood Chemistry	9
3.5 Pathology	10
3.6 Reproduction Studies	10
3.7 Physiological Studies	11
3.8 Behavioral Studies	13
4 DISCUSSION	13
5 CONCLUSIONS	14
LITERATURE CITED	15
APPENDIXES	17
A. Chemical Analysis of White Phosphorus Smoke	17
B. Details of Toxicological Evaluation	19
C. Pathology Report - Spontaneous Deaths	39
D. Pathology Report - 6-Week Exposure	55
E. Pathology Report - 13-Week Exposure	79
F. Pathology Report - 13-Week Exposure and 4-Week Recovery Period	103
DISTRIBUTION LIST	127

5/6

THE SUBCHRONIC EFFECTS OF REPEATED EXPOSURE TO WHITE PHOSPHORUS/FELT SCREENING SMOKE IN RATS

1. INTRODUCTION

Toxicity studies were conducted to determine the subchronic effects of repeated exposure to three concentration levels of a white phosphorus screening smoke. This smoke, one of the candidate screening smokes, was generated by burning pieces of white phosphorus-impregnated felt.

The following studies were performed to obtain a subchronic toxicity profile on rats: inhalation toxicity of multiple exposures, including pathology, hematology, and blood chemistry, and effects of exposures on reproduction, behavior, and physiology.

2. EXPERIMENTAL PROCEDURES

2.1 Chamber operation and sample collection.

Animals for the inhalation studies were placed in compartmented stainless-steel wire-mesh cages. The cages were put on racks in a 20-cubic-meter exposure chamber.

Military grade white phosphorus, which had been forced under pressure into thick pieces of wool felt, was cut into cubes having specific weights. For each exposure, one of these cubes on an aluminum foil pan of a known weight was placed on an electric hotplate within the chamber. The hotplate was a fast-heating unit capable of temperatures in excess of 700°F.

The chamber door was closed and sealed, and the chamber exhaust system was shut down. The hotplate was plugged in and the white phosphorus/felt ignited, usually in less than 2 minutes, producing a dense white smoke. This smoke is made up of a number of oxides of phosphorus: $P_4O \cdot P_2O \cdot P_2O_3$ (or P_4O_6), PO_2 (or P_2O_4), P_2O_5 (or P_4O_{10}), and P_2O_6 ; but only three of these species - P_2O_3 , PO_2 , and P_2O_5 - are well established.¹ All three of these oxides are deliquescent and they could, therefore, contribute various aqueous droplets to the smoke. The reaction of P_2O_5 with atmospheric moisture has already been described; the corresponding reaction for P_2O_3 produces o-phosphorus acid. The tetraoxide, P_2O_4 , reacts with water to produce equimolar solutions of o-phosphorous and o-phosphoric acid.¹ The white phosphorus/felt burning time ranged from less than 1 minute for some of the small cubes to over 9 minutes for one of the large cubes. The smoke was contained within the chamber for 15 minutes before being removed by the exhaust system.

Smoke sampling started after 7 minutes of exposure. The samples were collected on Gelman Type-A glass fiber filter pads. Twelve liters of chamber air, at the rate of 3 liters per minute, were drawn through the filter pads contained in the air sampler probe. The filter pads were placed in Erlenmeyer flasks. Particulate material was extracted with distilled water, boiled for 10 minutes, and titrated for phosphoric acid content (see appendix A).

Air samples were also collected in a modified Rochester cascade impactor and analyzed for phosphoric acid content, and the mass median diameter of the airborne particles was derived.

2.2 Animals used.

Four groups of Sprague-Dawley rats were used, one for each dose level plus the controls. Thirty-six male and 36 female rats were used for the high- and intermediate-exposure levels with 18 rats of each sex acting as controls for each level. Only 18 males and a like number of females were exposed to the low-dose level. Nine animals of each sex acted as controls for this exposure.

Additional groups of rats were exposed and the rats were rated on different schedules to determine the effects of repeated exposures on mammalian germ cells and reproductive performance. Procedures used are discussed in a previous report.*

2.3 Animal exposures and observations.

Animals were exposed for 15 minutes daily, 5 days a week for 13 weeks. The rats were observed for toxic signs before and after each exposure. At regular intervals during the 13-week exposure and 4-week postexposure periods, all animals were weighed.

After 6 weeks, one-third of the animals were withdrawn from exposure. Half of these animals were bled and submitted to pathology. The other half were examined to see if any physiological or behavioral changes occurred. When the 13 weeks of exposure ended, the remaining animals were separated into two groups. One group was divided as were the 6-week animals with half going to pathology and half for physiological and behavioral studies. The remaining animals were held for a 4-week postexposure period to observe any recovery that might occur. Then they were bled and submitted for pathological evaluation.

2.4 Blood studies.

Blood samples in rats were analyzed for triglycerides, cholesterol, glucose, urea nitrogen, creatinine, uric acid, sodium, potassium, chloride, carbon dioxide, alkaline phosphatase, albumin, globulin, calcium, phosphorus, serum glutamic pyruvic transaminase, serum glutamic oxalacetic transaminase, red blood cell count, white blood cell count, differential white blood cell count, hematocrit, and hemoglobin.

2.5 Pathology.

Gross and microscopic pathological examinations of the following body organs were performed for all animals submitted: heart, lung, liver, spleen, kidney, brain, eye, trachea, nasal turbinate, adrenal, stomach, urinary bladder, pancreas, thyroid, esophagus, duodenum, colon, lymph node, thymus, testis, epididymus, ovary, uterus, bone marrow, and skin. In addition to those animals sacrificed at 6, 13, and 17 weeks, animals that died during the experiments were submitted for necropsies. The number sacrificed at each time interval was adjusted according to the number of animals that died during that interval.

*Starke, W. C., and Pellerin, R. J. Technical Memorandum. The Effects of CS-2 on Reproduction in Rats (in preparation). 1980.

2.6 Physiological and behavioral studies.

Animals were submitted for physiological and behavioral evaluations using the procedures described in a previous publication.²

3. **RESULTS**

3.1 Exposure conditions.

Target concentrations were 1000, 500, and 200 mg/m³. Target and actual concentrations are used interchangeably in the appendixes. Concentrations are also referred to as high, intermediate, and low, respectively. To obtain desired concentrations, 20-, 10-, or 3-gram cubes of the white phosphorus-impregnated felt were used. When ignited, a dense white smoke-like mist was produced. The droplets of this mist had a mass median diameter of 0.5 micron. A charred residue weighing approximately 27% of the original weight remained after burning.

Appendix B, table B-1, shows the mean exposure doses produced by each cube recorded.

3.2 Toxic signs.

Animals were not visible for observation until the smoke cloud was evacuated from the chamber. Visible toxic signs were not observed during the first 3 days of exposure to the high dose. One hour after the fourth exposure one female rat died. Two other rats wheezed for about 2 hours. Of the 72 rats starting this high-dose study, 23 died during the first 6 weeks of exposure. Four more died during the remaining 7 weeks. Transitory dyspnea and wheezing were observed in many rats, with recovery within 2 hours of each exposure. The cumulative exposures at which deaths occurred are shown in appendix B, table B-2.

Rats exposed to the intermediate and low doses showed no visible toxic signs. None of these animals died.

3.3 Body-organ weights.

There were no consistent changes in total body weights or in weights of the organs regardless of dose or time. Means and standard errors for body and organ weights of control and exposed rats are shown in tables B-3 and B-4, appendix B. Significant differences, where they did occur, are indicated.

3.4 Hematology and blood chemistry.

No agent-related changes were observed in the blood chemistry and hematology analyses. "T" test evaluations indicated there were no significant agent-related differences between control and exposed values. No sex-related changes were observed. Means and standard errors of all blood constituents analyzed are shown in tables B-5 through B-10, appendix B.

3.5 Pathology.

3.5.1 Spontaneous deaths.

A high mortality rate (about 40%) occurred in the group of colony rats exposed to the 1161 mg/m³ dose level. The deaths appear to be agent and dose related. None of the rats exposed to the intermediate- or low-dose levels died. A complete description of pathological findings is presented in appendix C.

3.5.2 Six-week exposure.

Although laryngitis or tracheitis was not observed in any of the control animals, all laryngeal and tracheal specimens from rats exposed to the high-dose level displayed a moderate-to-severe laryngitis/tracheitis. Fifty percent of the rats receiving the intermediate-dose level displayed a minimal-to-mild tracheitis while one-third had a mild laryngitis. Only one rat receiving the low dose displayed tracheitis. Four of six high-dose rats had a minimal-to-severe interstitial pneumonia. Two of the 18 control rats displayed a minimal interstitial pneumonia. A complete description of these pathological findings is presented in appendix D.

3.5.3 Thirteen-week exposure.

Although none of the control rats displayed laryngitis or tracheitis, all of the male rats receiving the high-dosage level of white phosphorus/felt exhibited a moderate laryngitis. Of the female rats receiving the same dosage level (1161 mg/m³) of white phosphorus/felt, one of two larynges examined displayed a moderate laryngitis; whereas two out of three tracheae displayed mild-to-moderate tracheitis. Three out of six female rats receiving the medium-dosage level of white phosphorus/ felt displayed a moderate tracheitis; whereas three out of five male rats receiving the same level exposure displayed slight-to-moderate tracheitis. None of the low-dosage-level animals displayed laryngitis or tracheitis. A complete description of these pathological findings is presented in appendix E.

3.5.4 Thirteen-week exposure with four-week recovery period.

Significant findings were limited to the respiratory tracts of the rats exposed to the high- and intermediate-dose levels. Lesions were noted in the larynx or trachea of 15 of 16 high-dose rats and 20 of 24 intermediate-dose rats. Pulmonary lesions were noted in 11 of 16 high-dose and 6 of 24 intermediate-dose rats. None of the control or low-dose animals exhibited significant lesions. A complete description of these pathological findings is presented in appendix F.

3.6 Reproduction studies.

To assess the effects of white phosphorus/felt smoke on reproduction processes in rats, three studies were conducted: teratology, dominant lethal mutation, and reproduction in a single generation. Each of these studies was conducted at both the high- and intermediate-exposure levels and in the control (smoke-free) atmosphere.

At the high-dose level of each study, several of the animals died. Five of the 24 high-dose teratology females died before their scheduled necropsy date. Twelve of the 20 males for the single-generation study died prior to their time for mating. Nine of the 20 males for the dominant lethal mutation study died before they could be mated.

3.6.1 Teratology.

Pregnant females were exposed to test and control atmospheres from days 6 to 15 of gestation. There were no major abnormalities found among the control or test fetuses. The pregnancy rates for control, intermediate-dose and high-dose smoke were 100%, 100%, and 90%, respectively. Data on the number and condition of implants, the mean body weights of the fetuses, and the averaged data on implants appear in appendix B, table B-11. Statistical analysis of these data showed no differences between control and treatment groups.

3.6.2 Dominant lethal mutation.

In the dominant lethal mutation study, analysis by the chi-square method showed that, for the first mating, significantly more of the dams mated to intermediate-dose males had one or more resorptions as compared to the dams mated to control males, appendix B, table B-12. This difference was not observed at the high dose, indicating that the difference observed with the intermediate-dose group was probably not attributable to the effects of white phosphorus/felt smoke.

3.6.3 Single-generation reproduction.

The single-generation reproduction study results are more difficult to analyze. Although there were no significant differences in the body weights of the pups at birth, survivability of the exposed pups was significantly lower than that of the control pups, appendix B, table B-13. Exposures continued for the pups and dams through this 21-day period. Weight gain in the surviving exposed pups was less than that of the control pups, appendix B, table B-14. The difficulty arises from the inability to discern whether the deficiency in weight gain was (1) the result of the exposed dams not caring for their pups, (2) a decrease in mammary gland secretion, (3) the inability of the pups to nurse because of irritated and congested tracheas, or (4) exposure effects.

3.7 Physiological studies.

Physiological measurements were made on days 3 and 4 after the animals were removed from the exposure. The number of animals (sample size) used in the tests are shown in appendix B, tables B-15 and B-16.

An analysis of variance (ANOVA) was made on the data to determine the effect of dose (exposure level), sex difference, and the effect of dose within sexes. Following this, a T-test was performed to locate any significant differences.

The logic for determining a physiological effect from the white phosphorus/felt smoke required that the following conditions apply: (1) a significant difference at a $P \leq 0.05$ level be evidenced by the analysis of variance and Student T test, (2) the differences must be dose related and directional so that significant differences occurring in a low-dose group that were not reinforced by similar effects in the high-dose groups would be considered a chance occurrence, and (3) the difference should be time related unless an adaptive response for increased tolerance has developed. With a bioaccumulated material, it would be expected that effects shown after a 6-week exposure would be repeated after a 13-week exposure. With white phosphorus smoke, it is interesting to note from pathology (page 10) that upper airway irritation, i.e., tracheitis and laryngitis,

decreased after a 6-weeks' exposure. Assuming that phosphorous pentoxide unites with water lining the respiratory tract to form a phosphoric acid, the response to this irritant may be decreasing with time of exposure and consequently the application of present statistical judgments takes the possibility of increased tolerance into effect.

The results are shown in appendix B, tables B-15 through B-17. At both 6- and 13-weeks' exposure, significant sex differences were shown which appear to be related to the greater size of the male rats. These included breathing volumes and the increased inhaled volume responses to 6% CO₂. In addition, blood pressures from male rats were slightly higher than those of female rats. This has been observed in other tests.

Some differences appeared to be the effect of dose and these are indicated in tables B-15 and B-16. However, none of these effects satisfied the logic described above for determining a physiological effect. The same applies for the sex-related doses. It is noted that the sample size in the high-dose exposures was reduced because of deaths during the procedure. It is possible that, if the sample size were larger for high-dose groups, a statistical significance would have been noted. It appears that some effect was developing in the group toward a reduction in tidal volume and an increase in breathing frequency with dose in the 13-week exposed animals. There is also indication that there was less growth in the male rats of this group.

Table B-17 contains the qualitative or nonparametric observations made on rats in the exposure groups. Three of the 13-week high-exposure groups had loud bubbly rales 3 days after exposure which were absent on the fifth post-exposure day.

Although no statistical validation for a functional effect exists, the presence of rales and the tendency for a reduction in tidal volume and an increase in breathing frequency in the high-dose animals indicate that a careful examination of the lung pathology data and the pulmonary resistance data be made.

The evaluation of the pulmonary function as a result of exposure to white phosphorus/felt smoke was carried out on rats that had been exposed for a period of 13 weeks. The method for estimating the pulmonary resistance in these animals was based on a comparison of the plethysmographic pressure during a respiratory cycle and the peak respiratory flow. The rationale for using this method is based on the principle that the changes in pressure in a body plethysmograph during a respiratory cycle are proportional to the alveolar pressure during post cycle. (Pulmonary resistance is determined by the ratio of alveolar pressure to the flow rate during a respiratory cycle; therefore, an estimate of the resistance can be determined by substituting the plethysmographic pressure for the alveolar pressure.)

The results of the pulmonary tests in the unanesthetized rat following 13-weeks exposure to white phosphorus/felt smoke are presented in appendix B, table B-18. There were no significant differences from control values in the respiratory rate or peak inspiratory flow rates in either males or females exposed to smoke. There was no significant difference in the estimated pulmonary resistance of the female rats following exposure. The estimated pulmonary resistance of exposed male rats had a tendency to increase slightly over that of the controls. This increase was significantly different in the low-dose rats but not in the high-dose animals; however, the indication is that some pulmonary damage may be present in the male rats exposed to white phosphorus/felt smoke. No rats were tested at 6 weeks postexposure.

3.8 Behavioral studies.

The behavioral responses of rats exposed to white phosphorus smoke for 6 or 13 weeks was measured using spontaneous activity and passive avoidance tests. In all instances, values are compared statistically to a group of unexposed control rats of equal age and weight.

3.8.1 Spontaneous activity.

The results of these tests are presented in appendix B, table B-19, which shows the gross and fine activity of the rats and the ratio between these two types of activity. Both male and female animals exposed for 6 weeks appeared to exhibit a difference in activity; however, the mean values presented were influenced by one or two animals from each group including controls. If the ratio is calculated from these means, the values become more uniform, indicating that spontaneous activity was not altered as a result of exposure.

3.8.2 Passive avoidance.

Passive avoidance test results are presented in appendix B, table B-20. The exposed female rats, at 1161- and 589-mg/m³ dose levels and 6-week/13-week exposure time periods, were no different from control animals. This indicated that the animals' ability to passively avoid shocks was not adversely affected by exposure to white phosphorus smoke. Male rats exposed to the high dose received more shocks than the control rats but statistically the values were not significantly different. The number of passive avoidance responses were no different. One of the 13-week high-dose male rats received an unusually large number of shocks with no attempted avoidance, raising the mean value for this group of animals. This animal had either a higher pain threshold than the other animals or a delay in determining how to avoid the shock. The lower-dose male rats showed no differences from the control animals.

4. DISCUSSION

When white phosphorus burns, it forms a number of oxides of phosphorus which are rapidly converted by moisture to phosphorus and phosphonic acids.^{1,3} Although at ordinary expected field concentrations (100-200 mg/m³), this may not be hazardous, some irritation to the eyes, nose, and throat may occur.⁴ Cullumbine⁵ exposed a total of 108 men to white phosphorus smoke at concentrations from 87 to 1770 mg/m³. Throat irritation was produced at all concentrations. From these data, it has been estimated that the minimum harassing concentration (requiring a respirator) is about 700 mg/m³. In the documentation for threshold limit values (TLV), it has been reported that concentrations of 100 mg/m³ were unendurable except to hardened workers.⁶ A TLV of 1 mg/m³ was established as the level to which nearly all workers may be repeatedly exposed without adverse effect. A man exposed to 100 mg/m³ would probably mask or remove himself from the smoke cloud.

In the study just completed, the 1000 mg/m³ dose, with 40% spontaneous deaths and injury to all other animals, was obviously a hazardous concentration. None of the animals exposed at the 500- or 200-mg/m³ dose levels died. The intermediate dose did, however, produce tracheitis in 50% of the rats. Only one of the low-dose animals showed any signs of irritation.

5. CONCLUSIONS

It would appear that the chance of injury is high at 1161 mg/m^3 and low at 193 mg/m^3 . At 589 mg/m^3 , one-half of the rats showed some exposure effects.

The rats apparently developed a tolerance to repeated exposures, particularly at the lowest exposure level.

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APPENDIX A

CHEMICAL ANALYSIS OF WHITE PHOSPHORUS SMOKE*

Exposure chamber air samples received were diluted with distilled water, shaken, then boiled 10 minutes to convert the phosphorus acids to orthophosphoric acid, then cooled to room temperature. Sample volume was measured after boiling. Using a pH meter as an indicator, the acid content was determined by titration against 0.10043 N sodium hydroxide to a pH of 9.6. After measuring the amount of 0.10043 N sodium hydroxide required to adjust the sample pH to 9.6, the following formula was used to determine normality of the orthophosphoric acid:

$$1. \quad N_{\text{sample}} \times V_{\text{sample}} = N_{\text{NaOH}} \times V_{\text{NaOH}}$$

N = Normality

V = Volume of sample in milliliters

The normality of the acid is multiplied by 32.66 (milligrams per milligram-equivalents of orthophosphoric acid).

$$2. \quad \frac{\text{milligram equivalents}}{\text{milliliters} \\ (\text{normality of sample})} \times \frac{32.66 \text{ milligrams}}{\text{milligram equivalents}} = \frac{\text{milligrams}}{\text{milliliters of} \\ \text{orthophosphoric acid}}$$

$$3. \quad \frac{\text{milligrams}}{\text{milliliters}} \times \text{milliliters} = \text{milligrams of orthophosphoric acid in sample}$$

* B. P. Pearce. Porton Technical Paper 154. The Stability of Red Phosphorus Compositions. June 1974. UNCLASSIFIED Paper.

APPENDIX B
DETAILS OF TOXICOLOGICAL EVALUATION

Table B-1. Inhalation Exposure Doses Produced by Burning Cubes of White Phosphorus/Felt

Cube weight gm	Average exposure concentration mg/m ³	Average daily 15-minute exposure Ct mg min/m ³	Total cumulative exposure Ct mg min/m ³
20	1161	17,415	1,097,151 (63 days)
10	589	8,833	556,480 (63 days)
3	192.5	2,887	178,991 (62 days)

Table B-2. Spontaneous Deaths in Rats Exposed to the High-Dose Concentration of White Phosphorus Smoke

Exposure day	Deaths		Cumulative exposure Ct mg min/m ³
	Male	Female	
4		1	54,225
5	1		69,225
9	1		143,975
10	1		158,820
11		2	176,340
12	1	3	194,790
13	1	1	210,960
14	2	1	225,120
16		1	251,925
23	2		376,260
24		1	397,650
25	1		420,330
26	1		441,300
27	1		460,545
29		1	499,560
34	1		588,045
47	1		813,021
54	1		936,820
57	1		995,426

Appendix B

Table B-3. Organ and Total Body Weights in Male Sprague-Dawley Rats After Repeated Exposures to White Phosphorus Smoke

Dose level	Exposure time	Number of animals	Total body		Heart		Lungs		Liver		Kidneys		Gonads	
			Mean	Standard error	Mean	Standard error	Mean	Standard error	Mean	Standard error	Mean	Standard error	Mean	Standard error
High	6 weeks	0	—	—	—	—	—	—	—	—	—	—	—	—
High	13 weeks	6	373	19	1.62	0.23	3.32+	0.22	14.9	1.8	2.78++	0.16	3.47	0.11
High	17++	9	455	30	1.77	0.20	3.55	0.35	17.2	3.4	3.12	0.45	3.40	0.60
control	6 weeks	3	382	26	1.50	0.14	3.60	0.14	18.2	1.4	2.93	0.47	3.67	0.41
control	13 weeks	3	389	14	1.53	0.17	2.93	0.12	15.2	2.9	3.00	0.82	3.67	0.19
control	17++	6	417	46	1.57	0.14	3.32	0.42	16.0	2.3	2.93	0.44	3.37	0.14
Intermediate	6 weeks	6	374	20	1.47	0.24	3.32	0.56	16.3	1.2	2.67	0.28	3.47	0.26
Intermediate	13 weeks	6	433	27	1.68	0.19	3.25	0.22	16.2	2.6	2.83	0.31	3.52	0.21
Intermediate	17++	12	461++	46	1.64	0.21	3.36	0.37	17.9	3.1	3.11	0.31	3.62	0.47
Intermediate	6 weeks	3	368	15	1.37	0.09	3.07	0.09	15.4	0.8	2.93	0.26	3.27	0.20
control	13 weeks	3	456	34	1.63	0.17	3.27	0.17	16.0	1.2	2.87	0.05	3.30	0.33
control	17++	6	509	32	1.77	0.12	4.23	0.82	19.5	3.6	3.52	0.36	3.02	1.36
Low	6 weeks	6	346	20	1.50	0.15	3.50	0.36	16.2	1.5	2.67	0.11	3.35	0.10
Low	13 weeks	6	422	18	1.43	0.20	3.30	0.36	15.9	1.5	2.75	0.21	3.45	0.18
Low	17++	6	453	55	1.48	0.23	2.98++	0.38	17.8	2.5	2.78	0.35	3.48	0.37
Low	6 weeks	3	338	53	1.43	0.17	3.35	0.15	15.3	3.4	2.73	0.33	3.13	0.10
control	13 weeks	3	444	63	1.53	0.19	3.43	0.25	18.8	2.8	3.00	0.29	3.63	0.25
control	17++	3	484	11	1.90	0.24	3.43	0.12	18.5	3.0	3.30	0.24	3.57	0.17

+ - Significantly higher than control based on statistical "t" test.

++ - Significantly lower than control based on statistical "t" test.

+++ - Four weeks' postexposure.

Appendix B

Table B-4. Organ and Total Body Weights in Female Sprague-Dawley Rats After Repeated Exposures to White Phosphorus Smoke

Dose level	Exposure time	Number of animals	Total body		Heart		Lungs		Liver		Kidneys		Gonads	
			Mean	Standard error	Mean	Standard error	Mean	Standard error	Mean	Standard error	Mean	Standard error	Mean	Standard error
weeks														
High	6	6	241	12	0.93	0.09	2.63+	0.38	9.0	0.6	1.65	0.13	0.200	0.082
	13	6	241	19	1.15	0.10	2.67	0.41	9.4	1.9	1.82	0.31	0.183	0.037
	17	7	255	16	1.06	0.14	2.76	0.57	10.1	0.7	1.79	0.16	0.143	0.050
High control	6	3	235	9	0.97	0.05	2.10	0.08	10.0	0.5	1.70	0.08	0.167	0.047
	13	3	270	12	1.10	0.14	2.43	0.19	9.4	1.2	1.87	0.12	0.200	-
	17	6	254	19	1.07	0.09	2.32	0.26	10.0	1.8	1.75	0.24	0.167	0.047
Intermediate	6	6	247	22	0.97	0.09	2.47	0.24	9.7	1.2	1.85	0.30	0.183	0.087
	13	6	269	21	1.20	0.13	2.75	0.25	10.3	1.1	1.90	0.21	0.200	0.037
	17	12	266	24	0.99++	0.10	2.41	0.18	9.4++	1.2	1.89++	0.18	0.150	0.050
Intermediate control	6	3	246	6	1.03	0.04	2.43	0.09	10.0	0.4	1.70	0.08	0.167	0.04
	13	3	260	28	1.17	0.12	2.50	0.08	9.9	0.6	1.83	0.09	0.200	-
	17	6	275	13	1.15	0.08	2.33	0.12	10.8	1.1	2.15	0.19	0.150	0.05
Low	6	6	232	16	1.08	0.11	2.48	0.23	9.8	1.2	1.90	0.20	0.167	0.047
	13	6	240	15	1.05	0.13	2.33	0.09	8.8	0.9	1.83	0.17	0.167	0.047
	17	6	287	23	1.08	0.11	2.45	0.13	10.7	1.4	1.98	0.15	0.183+	0.037
Low control	6	3	229	24	1.07	0.17	2.47	0.45	8.9	1.4	1.90	0.16	0.200	0.048
	13	3	243	20	1.07	0.05	2.37	0.05	9.3	0.8	1.93	0.20	0.233	0.05
	17	2	270	5	1.25	0.05	2.35	0.15	9.9	-	2.65	0.35	0.100	-

+ - Significantly higher than control based on statistical "t" test.

++ - Significantly lower than control based on statistical "t" test.

Table B-5. Hematology in Rats Exposed for Six Weeks to White Phosphorus Smoke

Cumulative exposure Ct mg min/m ³	Red blood cells			White blood cells			Hematocrit			Hemoglobin			Neutrophils		
	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation
499.560	6	6.32	0.77	6	8333.33	1129.40	6	35.70	4.46	6	13.53	1.59	-	-	-
Control	6	6.90	0.58	6	9650.00	1698.77	6	38.53	2.95	6	15.10	0.87	-	-	-
246.285	12	7.02	0.53	12	6633.33	1818.12	12	39.11	3.34	12	14.42	0.94	12	7.42	5.63
Control	6	6.90	0.34	6	5966.67	1188.37	6	38.10	2.19	6	14.18	0.84	6	7.06	4.16
84.019	12	6.50	0.54	12	8325.00	1799.13	12	35.68*	3.08	12	13.51*	0.94	12	8.92	5.25
Control	6	6.95	0.41	6	10233.33	1569.15	6	38.58	2.10	6	14.68	0.71	6	10.17	3.80

Cumulative exposure Ct mg min/m ³	Band cells			Lymphocytes			Monocytes			Eosinophils			Basophils		
	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation
499.560	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Control	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
246.285	-	-	-	12	91.50	5.06	-	-	-	6	2.17	1.67	-	-	-
Control	-	-	-	6	92.67	4.57	1	1.00	0.00	1	0.00	0.00	-	-	-
84.019	-	-	-	12	90.25	5.55	1	1.00	0.00	6	1.50	0.76	-	-	-
Control	1	1.00	0.00	6	88.00	4.93	3	2.00	0.82	2	2.00	1.00	-	-	-

* Significantly lower than control based on statistical "t" test.

NOTE: Statistical evaluation of differential-white count components, monocytes, eosinophils, basophils and band cells may be meaningless since normal occurrences range from 1% to 10%.

Table B-6. Hematology in Rats Exposed for Thirteen Weeks to White Phosphorus Smoke

Cumulative exposure Ct mg min/m ³	Red blood cells				White blood cells				Hematocrit				Hemoglobin				Neutrophils			
	No. of animals	Mean		Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	
		x 10 ⁶ /mm ³	x 10 ⁶ /mm ³	x 10 ⁶ /mm ³									ng/100 ml	ng/100 ml	%	%	%	%	%	
1,097,151	12	7.32*	0.59	12	15,125.00	6301.47	12	40.12†	3.40	1.2	15.43*	1.28	1.0	31.10**	12.04					
Control	6	8.12	3.46	6	14,350.00	3114.35	6	44.83	1.64	6	17.48	0.77	6	15.00	6.53					
556,480	12	7.13	0.41	12	7691.67*	607.53	12	39.05	2.26	12	14.77	1.22	12	15.67	12.47					
Control	6	7.38	0.68	6	8633.33	727.25	6	39.47	3.16	6	15.10	1.23	4	11.00	8.77					
178,991	12	7.04**	0.88	12	9533.33	2489.76	12	38.51†	4.63	12	13.95**	1.90	12	14.58	4.61					
Control	6	5.46	1.05	6	8416.67	2625.15	5	29.88	6.50	6	10.98	2.00	6	10.67	4.85					

Cumulative exposure Ct mg min/m ³	Band cells				Lymphocytes				Monocytes				Eosinophils				Basophils			
	No. of animals	Mean		Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	
		%	%	%																
1,097,151	-	-	-	-	10	66.70	13.10	1	1.00	0.00	6	3.50	1.38	-	-	-	-	-	-	
Control	-	-	-	-	6	86.33	7.23	-	-	-	2	2.00	1.00	-	-	-	-	-	-	
556,480	-	-	-	-	12	83.67	12.39	-	-	-	3	2.67	0.94	-	-	-	-	-	-	
Control	-	-	-	-	4	88.50	8.99	-	-	-	1	2.00	0.00	-	-	-	-	-	-	
178,991	-	-	-	-	12	84.67	4.48	3	1.33	0.47	4	1.25	0.43	-	-	-	-	-	-	
Control	-	-	-	-	6	88.83	4.78	-	-	-	2	1.50	0.50	-	-	-	-	-	-	

Table B-7. Hematology in Rats 30 Days After 13 Weeks' Exposure to White Phosphorus Smoke

Cumulative exposure Ct mg min/m ³	Red blood cells			White blood cells			Hematocrit			Hemoglobin			Neutrophils		
	No. of animals	Standard deviation	No. of animals	No. of animals	Standard deviation	No. of animals	Standard deviation	No. of animals	Standard deviation	No. of animals	Standard deviation	No. of animals	Standard deviation	No. of animals	Standard deviation
1,097;151	16	7.15	1.0	16	11,506.25	1,747.31	16	40.81	4.55	16	14.48	1.07	16	33.44 ⁺	16.56
control	12	7.29	0.7	12	9,558.33	2,457.80	12	39.31	4.44	12	14.45	1.84	12	20.08	11.16
556;480	24	7.07	0.57	24	8,854.17	1,918.55	24	38.57	3.00	24	14.44	0.99	24	11.17	6.38
control	12	7.47	0.43	12	9,575.00	2,876.09	12	44.06	2.39	12	15.02	0.92	12	13.92	5.12
178;991	12	6.67	0.93	12	10,958.33	1,141.97	12	37.74	4.94	12	14.96	1.86	12	12.42	5.33
control	5	6.61	0.70	5	11,600.00	1,404.28	5	36.96	3.91	5	14.74	1.22	5	12.90	5.60

Cumulative exposure Ct mg min/m ³	Band cells			Lymphocytes			Monocytes			Eosinophils			Basophils		
	No. of animals	Standard deviation	No. of animals	No. of animals	Standard deviation										
-	%	%	16	65.75	16.74	2	1.00	0.00	4	2.75	0.83	-	-	-	-
1	1.00	0.00	2	78.67	11.01	3	1.00	0.00	7	1.57	0.73	-	-	-	-
-	-	-	24	88.04	6.26	1	4.00	0.00	13	1.15	0.36	-	-	-	-
-	-	-	12	85.58	5.19	-	-	-	5	1.20	0.40	-	-	-	-
-	-	-	12	87.00	4.93	-	-	-	4	1.75	0.83	-	-	-	-
-	-	-	5	87.20	6.52	-	-	-	3	1.33	0.47	-	-	-	-

+ Significantly higher than control based on statistical "t" test.

++ Significantly lower than control based on statistical "t" test.

NOTE: Statistical evaluation of differential-white count components, monocytes, eosinophils, basophils, and band cells may be meaningless since normal occurrences range from 1% to 10%.

Table B-8. Blood Chemistry in Rats Blood After 6 Weeks' Exposure to White Phosphorus Smoke

Cumulative exposure C _t mg min/m ³	Glucose						Blood urea nitrogen						Creatinine						Sodium						Potassium					
	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation			
		mg/dl	mg/dl		mg/dl	mg/dl		mg/dl	mg/dl		mg/dl	mg/dl		mg/dl	mg/dl		mg/dl	mg/dl		mg/dl	mg/dl		mg/dl	mg/dl		mg/dl	mg/dl			
499.560	6	150.17*	12.40	6	23.00	3.06	6	0.70	0.08	6	145.17	1.77	6	4.43	0.55	6	4.43	1.02	6	4.97	0.90	6	4.52	0.31	6	4.47	0.61			
Control	6	187.17	30.10	5	22.83	2.07	6	0.63	0.09	12	146.58	1.38	12	4.52	0.51	12	4.52	0.31	12	4.52	0.31	12	4.47	0.36	12	4.47	0.36			
246.285	12	165.63	15.46	12	21.33	1.84	12	0.54	0.12	6	142.23	1.25	6	4.25	0.25	11	4.25	0.25	11	4.25	0.25	11	4.25	0.25	11	4.25	0.25			
Control	6	171.67	19.88	6	23.83	2.97	5	0.55	0.17	11	145.27	2.18	6	4.05	0.35	6	4.05	0.35	6	4.05	0.35	6	4.05	0.35	6	4.05	0.35			
84.019	11	181.18**	22.39	13	20.64	2.38	11	0.55	0.08	11	145.83	1.21	6	4.05	0.35	6	4.05	0.35	6	4.05	0.35	6	4.05	0.35	6	4.05	0.35			
Control	6	159.17	4.03	6	19.33	0.75	6	0.50	0.10	6	145.83	1.21	6	4.05	0.35	6	4.05	0.35	6	4.05	0.35	6	4.05	0.35	6	4.05	0.35			
Cumulative exposure C _t mg min/m ³	Chloride						Carbon dioxide						Uric acid						Total protein						Albumin					
	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation			
		meq/l	meq/l		meq/l	meq/l		meq/l	meq/l		meq/l	meq/l		meq/dl	meq/dl		meq/dl	meq/dl		meq/dl	meq/dl		meq/dl	meq/dl		meq/dl	meq/dl			
499.560	6	103.33	1.11	6	21.67	4.31	6	0.85	0.68	6	5.58	0.42	6	3.03	0.32	6	3.30	0.15	6	3.33	0.12	6	3.16	0.15	6	3.16	0.15			
Control	6	101.57	1.37	6	22.00	3.83	6	1.09	0.69	6	6.07	0.23	6	3.30	0.15	6	3.30	0.15	6	3.33	0.12	6	3.16	0.15	6	3.16	0.15			
246.285	12	102.92	2.36	12	22.33	2.39	12	1.32	0.78	12	6.17	0.22	12	3.26	0.26	12	3.26	0.26	12	3.26	0.26	12	3.26	0.26	12	3.26	0.26			
Control	6	102.57	2.13	5	23.50	4.19	6	1.48	0.84	6	6.05	0.26	6	3.26	0.26	6	3.26	0.26	6	3.26	0.26	6	3.26	0.26	6	3.26	0.26			
84.019	11	103.55	2.23	11	19.09	5.81	11	1.47	1.37	11	5.54	0.53	11	3.15	0.24	11	3.15	0.24	11	3.15	0.24	11	3.15	0.24	11	3.15	0.24			
Control	6	104.90	2.90	6	22.00	4.62	6	0.73	0.26	6	5.87	0.43	6	3.15	0.24	6	3.15	0.24	6	3.15	0.24	6	3.15	0.24	6	3.15	0.24			
Cumulative exposure C _t mg min/m ³	Globulin						Calcium						Phosphate						Cholesterol						Triglycerides					
	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation			
		g/m/dl	g/m/dl		g/m/dl	g/m/dl		g/m/dl	g/m/dl		g/m/dl	g/m/dl		mg/dl	mg/dl		mg/dl	mg/dl		mg/dl	mg/dl		mg/dl	mg/dl		mg/dl	mg/dl			
499.560	6	2.55	0.28	6	9.77	0.57	6	4.93*	0.65	6	75.00	6.11	6	33.67*	16.60	6	69.83	23.02	6	69.83	23.02	6	55.58	39.28	6	55.58	39.28			
Control	6	2.77	0.14	6	10.25	0.43	6	6.07	0.51	6	76.50	11.77	6	32.26	16.60	6	89.00	46.63	6	89.00	46.63	6	89.91	32.26	6	89.91	32.26			
246.285	12	2.73	0.16	12	9.79	0.36	12	5.93	0.54	12	73.17*	5.98	12	43.11	12.26	12	73.17*	5.98	12	73.17*	5.98	12	73.17*	5.98	12	73.17*	5.98			
Control	6	2.87	0.14	6	9.65	0.47	6	5.90	0.75	6	79.67	4.31	6	46.63	12.26	6	89.91	32.26	6	89.91	32.26	6	89.91	32.26	6	89.91	32.26			
84.019	11	2.57	0.18	11	9.67	0.33	11	5.35	1.13	11	106.18	9.84	11	51.83	24.13	11	106.18	9.84	11	106.18	9.84	11	106.18	9.84	11	106.18	9.84			
Control	6	2.72	0.27	6	9.87	0.25	6	6.15	0.52	6	97.67	9.84	6	51.83	24.13	6	106.18	9.84	6	106.18	9.84	6	106.18	9.84	6	106.18	9.84			
Cumulative exposure C _t mg min/m ³	Alkaline phosphatase						Glutamic transaminase						Glutamic pyruvic transaminase						Lactic dehydrogenase						Total bilirubin					
	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation			
		U	U		U	U		U	U		U	U		U	U		U	U		U	U		U	U		U	U		U	U
499.560	6	230.50	51.19	6	147.33	96.76	6	57.33	14.01	6	382.17	297.98	6	6.10	0.10	6	382.17	297.98	6	6.10	0.10	6	382.17	297.98	6	6.10	0.10			
Control	6	335.83	99.78	6	116.83	25.47	6	65.67	9.50	6	322.67	88.68	6	6.10	0.10	6	322.67	88.68	6	6.10	0.10	6	322.67	88.68	6	6.10	0.10			
246.285	12	366.17	139.84	12	120.83	54.22	12	53.08	10.85	12	287.50	173.36	12	6.10	0.10	12	287.50	173.36	12	6.10	0.10	12	287.50	173.36	12	6.10	0.10			
Control	6	301.00	96.39	6	140.00	81.76	6	59.17	11.99	6	147.27	113.08	6	6.10	0.10	6	147.27	113.08	6	6.10	0.10	6	147.27	113.08	6	6.10	0.10			
84.019	11	355.69	90.90	11	126.18	44.53	11	55.09	10.72	11	245.67	43.60	11	6.10	0.10	11	245.67	43.60	11	6.10	0.10	11	245.67	43.60	11	6.10	0.10			
Control	6	306.50	103.94	6	117.50	33.59	6	49.90	9.45	6	257.67	43.60	6	6.10	0.10	6	257.67	43.60	6	6.10	0.10	6	257.67	43.60	6	6.10	0.10			

* Significantly lower than control based on statistical "t" test.

** Significantly lower than control based on statistical "t" test.

Table B-9. Blood Chemistry in Rats Bled After 13 Weeks Exposure to White Phosphorus Smoke

Cumulative exposure Ct mg min/m ³	Glucose			Blood urea nitrogen			Creatinine			Sodium			Potassium		
	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation
499.560	12	178.58	22.76	12	20.58	4.33	12	0.57	0.07	12	146.61	2.05	12	4.43	0.65
Control	6	168.83	17.35	6	17.50	1.71	6	0.53	0.05	6	147.17	0.69	6	4.26	0.39
246.285	12	166.50	12.27	12	20.33	3.09	12	0.57	0.11	12	144.33	1.60	12	4.07	0.22
Control	6	174.50	14.85	6	21.33	3.14	6	0.63	0.5	6	144.50	0.98	6	4.20	0.15
84.019	12	171.00	16.87	12	21.67	1.49	12	0.52	0.09	12	145.72	0.86	12	4.15	0.43
Control	6	187.17	7.45	6	23.00	2.58	6	0.57	0.05	6	145.50	0.96	6	4.27	0.35

Cumulative exposure Ct mg min/m ³	Chloride			Carbon dioxide			Uric acid			Total protein			Albumin		
	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation
		meq/l	meq/l		meq/l	meq/l		mg/dl	mg/dl		gm/dl	gm/dl		gm/dl	gm/dl
12	106.00	2.68	1.2	12	17.58	4.31	12	1.33	1.38	12	5.85	0.27	12	3.30	0.11
6	107.50	0.50	6	1.71	1.71	6	0.81	0.81	6	5.75	0.32	6	3.30	0.14	
12	104.92	2.20	12	19.83	3.13	12	1.23	1.20	12	5.78	0.20	12	3.18	0.11	
6	104.67	3.45	6	21.17	3.18	6	1.18	0.64	6	5.90	0.19	6	3.33	0.09	
12	103.58	1.85	12	19.42	2.90	12	0.86	1.13	12	5.80	0.20	12	3.16	0.13	
6	104.33	1.97	6	19.00	2.89	6	1.08	0.41	6	5.83	0.25	6	3.10	0.16	

Cumulative exposure Ct mg min/m ³	Globulin			Calcium			Phosphate			Cholesterol			Triglycerides		
	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation
		gm/dl	gm/dl		gm/dl	gm/dl		mg/dl	mg/dl		mg/dl	mg/dl		mg/dl	mg/dl
12	2.55	0.13	12	9.68	0.57	12	4.92	1.09	12	111.42	30.22	12	36.25	54.69	
6	2.45	0.24	6	9.70	0.24	6	5.08	0.75	6	123.83	12.35	6	57.67	21.01	
12	2.60	0.15	12	9.37	0.23	12	4.70	0.45	12	74.25	9.69	12	31.33	33.24	
6	2.57	0.11	6	9.55	0.29	6	5.07	0.53	6	74.33	11.01	6	21.50	20.95	
:2	2.61	0.13	12	9.67	0.20	12	5.01	0.52	12	77.92	8.62	12	37.42	24.68	
6	2.73	0.19	6	9.43	0.30	6	5.02	0.49	6	77.33	6.05	6	20.67	18.12	

Cumulative exposure Ct mg min/m ³	Alkaline phosphatase			Glutamic oxaloacetic transaminase			Glutamic pyruvic transaminase			Lactic dehydrogenase			Total bilirubin		
	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation
		u.l.	u.l.		u.l.	u.l.		u.l.	u.l.		u.l.	u.l.		mg/dl	mg/dl
12	352.58**	103.00	12	109.08	37.23	12	55.17	13.40	12	299.00	168.56	12	0.10	0.00	
6	270.50	48.41	6	85.33	16.36	6	47.17	7.06	6	258.83	85.31	6	0.10	0.00	
12	262.00	72.63	12	91.67	38.87	12	45.75	10.30	12	290.50	117.40	12	0.16	0.05	
6	247.67	49.52	6	80.33	9.29	6	40.17	6.12	6	221.33	64.50	6	0.15	0.05	
12	257.42	101.75	12	108.50	30.82	12	56.92	20.54	12	253.67	49.95	12	0.11	0.03	
6	295.50	86.50	6	131.67	53.31	6	62.50	15.02	6	411.00	233.72	6	0.12	0.04	

* Significantly lower than control based on statistical "t" test.

** Significantly higher than control based on statistical "t" test.

Table B-10. Blood Chemistry in Pigs Placed 4 Weeks / After a 13-week Exposure to White Phosphorus Smoke

Cumulative exposure mg min/m ³	Glucose		Blood urea nitrogen		Creatinine		Sodium		Potassium			
	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation
499.560	16	173.81	12.53	14	22.25	2.22	16	0.72	0.09	16	147.19*	1.78
Control	12	175.17	20.32	12	22.67	3.77	12	0.69	0.08	12	145.92	1.04
286.285	24	180.21	27.87	24	20.04*	1.84	24	0.57	0.07	24	146.42	3.45
Control	12	168.08	15.26	12	21.58	1.32	12	0.59	0.13	12	147.50	1.55
84.019	12	196.75	26.54	12	20.33	1.31	12	0.53	0.06	12	145.58	1.66
Control	5	189.80	20.71	5	18.80	1.17	5	0.56	0.12	5	144.60	1.62

Cumulative exposure mg min/m ³	Chloride		Carbon dioxide		Uric acid		Total protein		Albumin			
	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation
16	105.56	1.17	16	19.31	2.52	16	0.98	0.39	16	5.86	0.25	16
12	105.83	2.37	12	21.17	2.82	12	1.02	0.70	12	5.92	0.33	12
24	103.42	2.18	24	20.92	2.71	24	1.01	0.67	24	5.64**	0.29	24
12	104.00	2.61	12	20.00	3.46	12	0.76	0.25	12	6.07	0.29	12
12	103.25	1.69	12	25.08	2.25	12	1.35	1.10	12	6.06	0.35	12
5	103.40	0.49	5	24.80	0.75	5	1.16	0.83	5	5.88	0.43	5

Cumulative exposure mg min/m ³	Globulin		Calcium		Phosphate		Cholesterol		Triglycerides			
	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation
16	2.58	0.19	16	9.41	0.27	16	4.69	0.63	16	79.37	18.26	16
12	2.59	0.11	12	9.42	0.49	12	4.32	0.61	12	78.33	19.72	12
24	2.47**	0.20	24	9.56	0.54	24	5.17	0.59	24	74.83*	6.54	24
12	2.67	0.12	12	9.75	0.42	12	5.01	0.67	12	84.75	11.55	12
12	2.67	0.19	12	10.10	0.24	12	5.33	0.56	12	94.83	6.87	12
5	2.60	0.28	5	9.98	0.55	5	5.36	0.13	5	75.60	9.44	5

Cumulative exposure mg min/m ³	Alkaline phosphatase		Glutamic oxaloacetic transaminase		Glutamic pyruvic transaminase		Lactic dehydrogenase		Total bilirubin			
	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation	No. of animals	Mean	Standard deviation
16	14	274.69	69.42	16	91.62	26.71	16	48.62	13.72	16	278.06	113.32
12	102.99	299.83	12	97.83	21.56	12	56.25	11.36	12	305.67	76.71	12
24	262.33	75.80	24	107.92	41.29	24	57.08	25.94	24	311.08	104.66	24
12	304.92	88.19	12	110.92	38.15	12	69.50	21.80	12	315.92	61.94	12
12	293.75	90.84	12	126.04	90.45	12	80.00	57.57	12	246.92	66.23	12
5	218.00	60.98	5	93.40	17.45	5	58.40	11.57	5	253.80	55.11	5

* Significantly lower than control based on statistical "t" test.

** Significantly higher than control based on statistical "t" test.

Table B-11. Teratologic Effects on Pups of Female Rats Futhanatized on Day 20 After Inhalation of White Phosphorus/Felt Smoke During Organogenesis

	Air controls	Dose level 589 mg/m ³	1,161 mg/m ³
Number pregnant	20	20	18
Percent pregnant	100	100	90
Total implants	238	242	221
Live implants	225	236	206
Dead implants	13	6	15
Percent dead implants	5.46	2.48	6.79
Mean weight in grams			
Male	3.91 ± 0.32	3.83 ± 0.36	3.87 ± 0.38
Female	3.76 ± 0.34	3.67 ± 0.30	3.66 ± 0.40
Mean body weight - all pups	3.84 ± 0.33	3.76 ± 0.34	3.77 ± 0.40
Average implants per pregnant female	11.90 ± 1.45	12.10 ± 1.37	12.28 ± 1.96
Average live implants per pregnant female	11.25 ± 1.41	11.80 ± 1.61	11.44 ± 2.28
Average dead implants per pregnant female	0.65 ± 0.67	0.30 ± 0.57	0.83 ± 1.10

Table B-12. Reproductive Data for Female Rats Mated to Males Exposed to White Phosphorus/Smoke
 (Numbers in parentheses indicate the number of animals in each category.)

Group	Week	Number mated	Number pregnant	M.I.*	C.L.I.	I.I.	P.I.I.I.	F.I.	R.I.	N.V.F./V.F.	N.V.F.≥1	N.V.F.≥2
Air control	1	20	14	70	(183)	(165)	(18)	(158)	(7)	7/158	5/9	2/12
	2	20	19	95	(253)	(238)	(15)	(223)	(15)	15/223	9/10	4/15
					13.32	12.53	0.79	11.74	0.79			
589 mg/m ³	1	20	19	95	(257)	(224)	(33)	(210)	(14)	14/210	13/6	1/18
	2	20	20	100	(278)	(259)	(19)	(249)	(10)	10/249	7/13	3/17
1161 mg/m ³	1	20	18	90	(256)	(226)	(30)	(215)	(11)	11/215	8/10	2/16
	2	20	16	90	(255)	(219)	(36)	(215)	(4)	4/215	4/14	0/18

*M.I. (matting index) = total number of females mated / total number of female mated

C.L.I. (corpus luteum index) = total number of corpora lutea / total number of pregnant females

I.I. (implantation index) = total number of implantation sites / total number of pregnant females

P.I.I.I. (postimplantation loss index) = total number of corpora lutea - total number of implantation sites / total number of pregnant females

F.I. (fetal index) = total number of viable fetuses / total number of pregnant female

R.I. (resorption index) = total number of deaths (early and late) / total number of pregnant female

N.V.F./V.F. = total number of observable fetuses / total number of viable fetuses

N.V.F.≥1 = total number of females with one or more observable fetuses

N.V.F.≥2 = total number of females with two or more observable fetuses

N.V.F.≥3 = total number of females with one or very observable fetuses

**Significant at p < 0.05.

Appendix B

Table B-13. Viability, Survival, and Lactation Indices in a Single-Generation Study of Reproduction Performance During Exposure to White Phosphorous/Felt Smoke

Index	Generation	Control	Low dose *	High dose **
Viability index	F1	99.48	93.23	64.12
Survival index: day 21	F1	98.44	92.19	30.23
Lactation index	F1	98.95	98.88	47.15

* Low dose animals exposed to 589 mg/m³ white phosphorus/felt smoke.
 ** High dose animals exposed to 1,161 mg/m³ white phosphorus/felt smoke.

Table B-14. Single-Generation Reproduction Study of White Phosphorus in Rat Smoke

Days Post-natal	Weight change (mean and SD)					
	Males			Females		
	Control	Low dose*	High dose**	Control	Low dose*	High dose**
1	(103) 6.63 ± 0.57	(90) 7.26 ± 0.74	(136) 6.38 ± 0.62	(89) 6.49 ± 0.69	(102) 6.76 ± 0.75	(166) 6.01 ± 0.69
4	(102) 10.41 ± 1.27	(84) 11.30 ± 1.15	(92) 9.41 ± 1.48	(89) 10.39 ± 1.54	(95) 10.71 ± 1.19	(101) 8.99 ± 1.59
7	(102) 15.36 ± 2.18	(84) 16.66 ± 1.67	(84) 13.97 ± 1.86	(89) 15.45 ± 2.69	(95) 15.64 ± 1.75	(90) 13.62 ± 2.26
14	(100) 29.59 ± 4.19	(83) 31.39 ± 3.52	(47) 26.46 ± 5.03	(89) 29.97 ± 5.31	(94) 29.46 ± 2.72	(44) 25.93 ± 5.53
21	(100) 45.05 ± 8.16	(83) 50.62 ± 5.34	(47) 41.10 ± 9.52	(89) 45.16 ± 9.71	(94) 46.55 ± 5.09	(44) 41.50 ± 11.15

* Low dose, 589 mg/m³.
 ** High dose, 1,161 mg/m³.

Table B-5. Physiological Effects of Short-Term Exposure to Various Concentrations of Inhalants After 6 Weeks' Exposure

Measurement	Time	Concen-	Exposure			High-dose			Significance of F Ratio			
			Low-dose			All			Dose			
			All	Males	Females	All	Males	Females	All	Males	Females	
Samples size			12	6	6	7	3	3	1	1	1	
Mean			54	35	76	59	37	76	300	54	347	10
Standard deviation			317	200	615	368	233	606	1007	63	1032	.05
Degrees of freedom, total			24	12	12	24	12	12	107	107	107	.00
Temperature, rectal	min		34	126	52	35	20	50	34	124	52	.07
Temperature, rectal	max		35	126	52	35	20	50	35	124	52	.01
Rectal volume	min		48	207	72	47	206	72	37	172	27	.0001
Rectal volume	max		51	212	75	50	212	75	37	172	27	.0001
Breathing frequency												
Breathing frequency	breaths/min		17	106	24	193	17	111	14	66	17	.04
Breathing frequency	per minute		107	60	175	107	60	175	107	60	175	.05
Respiratory rate, CO ₂	min		460	307	528	457	305	528	462	305	528	.0001
Respiratory rate, %CO ₂	min		459	305	527	460	303	527	464	303	527	.0001
Minute volume, %CO ₂	min		156	23	346	152	22	345	151	219	345	.05
Breathing frequency, %CO ₂	min		118	5	127	4	116	5	120	4	118	.05
Blood pressure, systolic												
Blood pressure, systolic	mm Hg		118	61	141	60	135	51	178	39	197	.35
Heart rate, from blood pressure	min		413	61	431	40	395	51	397	33	359	.37
Heart rate, from ECG	min		459	47	465	53	459	47	464	53	451	.01
ECG-P wave amplitude	mv		68	52	72	66	50	70	65	50	70	.027
P-wave duration	sec		0.017	0.022	0.017	0.017	0.017	0.017	0.017	0.017	0.017	.001
P-R interval	sec		0.046	0.093	0.046	0.044	0.046	0.046	0.046	0.044	0.046	.001
Q-R-S interval	sec		0.044	0.092	0.044	0.043	0.044	0.044	0.044	0.043	0.044	.001
Q-T interval	sec		0.062	0.134	0.062	0.064	0.062	0.064	0.062	0.064	0.062	.001
R-wave amplitude	mv		359	509	363	364	365	365	364	365	365	.05
R-wave amplitude	mm		32	35	30	34	32	34	32	34	32	.05
Median	min		12.5	3.0	11.3	5.0	9.3	11.3	10.4	5.7	7.3	.05

* Values indicate standard deviation.

† Student's *t*-test = different from control at P < 0.05.

Appendix B

Table B-16. Physiological Effects of White Phosphorus/Felt Smoke on Rats After 13 Weeks' Exposure

Measurement	Units	Exposure						ANOVA					
		Control		Low-dose		High-dose		Significance of F ratio (A)		Significance of F ratio (A)		Significance of F ratio (A)	
		All	Males	Females	All	Males	Females	All	Males	Females	Dose	Male x female	Dose: male
Sample size		12	6	6	12	6	6	7	4	4			
Weight	grams	365 ± 54	441 ± 16	288 ± 28	358 ± 109	458 ± 39	257 ± 19	324 ± 78	235 ± 25*	248 ± 30			
Temperature, rectal	degrees F	99.9 ± .54	99.9 ± .77	100.0 ± .17	100.8 ± .96*	100.1 ± .60	101.5 ± .68*	99.6 ± .28	99.7 ± .21	99.6 ± .46*			.0001
Total volume	ml	2.20 ± .39	2.51 ± .25	1.49 ± .22	1.78 ± .37**	2.06 ± .31**	1.50 ± .12*	1.86 ± .26	1.91 ± .17*	1.79 ± .43			.0001
Minute volume	ml	215 ± 62	264 ± 43	166 ± 30	185 ± 30	186 ± 34**	174 ± 24	215 ± 28	223 ± 16	207 ± 42			.0001
Breathing frequency	breaths per minute	97 ± 18	105 ± 12	88 ± 16	106 ± 17	95 ± 16	116 ± 11	117 ± 14**	116 ± 16	117 ± 13			.0001
Total volume, % CO ₂	ml	3.14 ± .48	3.50 ± .31	2.78 ± .31	2.80 ± .52	3.21 ± .37	2.59 ± .21*	3.06 ± .38	3.17 ± .35	2.91 ± .43			.0001
Breathing frequency, % CO ₂	ml/min	416 ± 93	467 ± 106	371 ± 46	386 ± 60	419 ± 47	358 ± 50	426 ± 83	470 ± 85	373 ± 41			.0001
Blood pressure, systolic	mm Hg	113 ± 22	133 ± 28	134 ± 16	141 ± 20	132 ± 23	149 ± 14	140 ± 21	149 ± 24	129 ± 13			.0001
Heart rate (from blood pressure)	b/min	119 ± 6.0	122 ± 7.1	117 ± 3.2	119 ± 7.6	123 ± 6.3	115 ± 6.4	119 ± 6.1	121 ± 7.4	117 ± 3.6			.0001
Heart rate (from ECG)	b/min	389 ± 30	383 ± 29	394 ± 33	456 ± 38	445 ± 34**	467 ± 44*	418 ± 29	417 ± 21	470 ± 44			.0001
ECG P-wave amplitude	mv	.055 ± .016	.060 ± .027	.050 ± .027	.043 ± .035	.075 ± .012	.090 ± .049	.059 ± .023	.040 ± .010	.047 ± .015			.0001
P-wave duration	sec	.016 ± .002	.017 ± .001	.016 ± .003	.017 ± .001	.018 ± .001	.017 ± .001	.017 ± .001	.017 ± .001	.017 ± .001			.0001
P-P interval	sec	.047 ± .006	.045 ± .004	.049 ± .007	.045 ± .003	.065 ± .002	.045 ± .004	.044 ± .002	.044 ± .002	.043 ± .002			.0001
O-R-S interval	sec	.016 ± .002	.016 ± .002	.016 ± .001	.016 ± .002	.017 ± .002	.015 ± .002	.016 ± .001	.016 ± .001	.016 ± .001			.0001
P-wave amplitude	mv	.053 ± .005	.052 ± .005	.053 ± .005	.055 ± .006	.054 ± .007	.055 ± .005	.055 ± .004	.055 ± .005	.054 ± .004			.0001
T-wave amplitude	mv	.12 ± .13	.25 ± .10	.23 ± .17	.36 ± .08	.34 ± .08	.38 ± .08	.24 ± .19	.37 ± .04	.07 ± .15			.0001
Peak-to-peak run	min	12.6 ± 4.2	20.3 ± 4.9	14.8 ± 1.8	12.8 ± 4.1	12.6 ± 3.8	12.9 ± 4.8	10.9 ± 3.5	11.7 ± 2.5	9.8 ± 5.0			.0001

*Values indicate standard deviation.

**Student's "t" test - different from control at P < .01.

***Student's "t" test - different from control at P < .05.

****Student's "t" test - different from control at P < .0001.

Table B-17. Qualitative Observations on Rats Exposed to White Phosphorous/Felt Smoke

Observation	Number responding					
	Exposure period					
	6 Weeks			13 Weeks		
	Control	Low dose	High dose	Control	Low dose	High dose
Vasomotor reflex	12/12	12/12	6/6	12/12	12/12	7/7
Light reflex	12/12	12/12	6/6	12/12	12/12	7/7
Respiratory response to CO ₂	12/12	12/12	6/6	12/12	12/12	7/7
Sniffing response	9/12	12/12	6/6	11/12	12/12	7/7
Moist rales	0/12	0/12	0/6	0/12	0/12	3/12

Appendix B

**Table B-18. Pulmonary Responses of Rats Exposed to White Phosphorus/Felt
Smoke for 13 Weeks**

Condition	Sex	Number of animals	Response (mean and SE)		
			Estimated pulmonary resistance	Respiratory rate	Peak inspiratory flow
			cmH ₂ O/l/sec	resp/min	ml/sec
Control	Male	5	1.03 ± 0.06	179 ± 14.3	24.8 ± 0.92
High dose	Male	4	1.29 ± 0.18	171 ± 2.74	24.7 ± 0.70
Low dose	Male	5	1.75 ± 0.27	142 ± 5.07	24.1 ± 0.55
Control	Female	6	1.58 ± 0.88	120 ± 3.3	22.6 ± 0.90
High dose	Female	3	1.75 ± 0.27	145 ± 26.6	24.2 ± 0.59
Low dose	Female	6	1.33 ± 0.13	134 ± 5.8	23.3 ± 0.72

Appendix B

Table B-19. The Spontaneous Activity Responses of Rats Exposed to White Phosphorus/Felt Smoke

Condition	Sex	Number of animals	Response (mean and SE)		Gross activity fine activity ratios	
			Gross activity	Fine activity	from individual data	from mean
6-Week exposure						
Control	Female	6	457 + 143	1680 + 512	5.12 + 1.39	3.68
High dose	Female	3	502 + 21	1739 + 140	3.49 + 0.37	3.46
Low dose	Female	6	574 + 134	1874 + 353	6.32 + 3.40	3.20
Control	Male	6	278 + 122	1093 + 327	5.10 + 0.53	3.93
High dose	Male	3	336 + 65	723 + 319	1.97 + 0.37	2.15
Low dose	Male	6	760 + 109	2253 + 306	3.07 + 0.34	2.96
13-Week exposure						
Control	Female	6	608 + 133	160 + 300	2.82 + 0.46	2.65
High dose	Female	3	462 + 38	1548 + 149	3.42 + 0.53	3.35
Low dose	Female	6	575 + 177	1652 + 421	3.44 + 0.46	2.87
Control	Male	6	601 + 138	1287 + 176	2.40 + 0.29	2.14
High dose	Male	4	420 + 48	1368 + 308	2.86 + 0.54	2.91
Low dose	Male	6	450 + 123	1651 + 191	3.25 + 0.23	3.67

Table B-20. The Passive Avoidance Responses of Rats Exposed to White Phosphorus/Felt Smoke

Condition	Sex	Number of animals	Number of passive avoidance responses (mean and SE)	Total number of shocks (mean and SE)	Time in shock (mean and SE)
<i>6-Week exposure</i>					
Control	Female	6	3.3 + 0.8	8.3 + 2.8	0.86 + 0.23
High dose	Female	3	4.7 + 1.7	8.7 + 5.6	0.84 + 0.52
Low dose	Female	6	3.0 + 1.0	7.0 + 1.9	0.73 + 0.19
Control	Male	6	2.7 + 0.5	6.2 + 1.9	0.72 + 0.25
High dose	Male	3	5.3 + 1.8	10.3 + 3.3	1.24 + 0.12
Low dose	Male	6	2.7 + 0.5	6.0 + 1.5	1.73 + 0.32
<i>13-Week exposure</i>					
Control	Female	6	3.7 + 0.8	9.3 + 3.2	0.87 + 0.35
High dose	Female	3	3.0 + 0.6	10.7 + 0.6	1.13 + 0.15
Low dose	Female	6	7.7 + 3.3	10.2 + 2.7	1.21 + 0.32
Control	Male	6	1.8 + 0.3	7.0 + 3.1	1.15 + 0.64
High dose	Male	4	3.8 + 1.2	25.5 + 20.9	2.33 + 1.89
Low dose	Male	6	2.3 + 0.6	5.3 + 2.4	0.55 + 0.27

Appendix B

APPENDIX C
PATHOLOGY REPORT - SPONTANEOUS DEATHS

PATHOLOGY REPORT
PROJECT SMOKE II, WHITE PHOSPHORUS/FELT
INHALATION STUDY - SPONTANEOUS DEATHS
"COLONY" RATS

I. INTRODUCTION.

Six and thirteen week studies designed to assess potential local and systemic toxic effects of inhalation exposure of the agent White Phosphorus/Felt were performed on Edgewood Area Colony Rats. Three dosage levels of the agent were tested. High (1000 mg/m^3) dosage level studies began on 31 July 1978; Medium (500 mg/m^3) dosage level studies began on 7 August 1978. Low (200 mg/m^3) dosage level studies began on 11 September 1978. All rats were approximately six weeks of age at the beginning of the studies and were housed in Bldg E3266.

Of forty-three rats (19 male, 24 female) receiving the high (1000 mg/m^3) dosage level exposure to White Phosphorus/Felt, twenty-nine (15 male, 14 female) died spontaneously during the studies while one control animal and none of the medium or low dose group animals died.

Following necropsy, tissues were imbedded in paraffin and subsequently processed for staining with hematoxylin and eosin. The following tissues were evaluated microscopically: nasal turbinates, larynx, trachea, lungs, heart, esophagus, stomach, small intestine, pancreas, large intestine, liver, adrenal, thyroid, thymus, kidney, bladder, ovary/teste, uterus, mammary gland, prostate, bone marrow, spleen, brain, eye and pituitary.

Histologic findings are tabulated in Tables 1-3. Since one or more tissues from various animals were lost at necropsy or during processing, one must calculate the incidence of lesions based upon the number of tissues examined rather than on the number of animals necropsied.

Two previous reports have been submitted regarding the histopathological findings in the colony rats that completed the six and thirteen week studies. This report deals only with the histopathological findings in those animals that died spontaneously before completing the studies.

II. RESULTS.

The microscopic observations are presented in the Histopathology Incidence Tables.

a. Table 1 tabulates incidence of lesions by organ observed in male and female rats dying spontaneously.

b. Table 2 tabulates incidence and severity of lesions by organ observed in male rats dying spontaneously from the high (1000 mg/m^3) dosage level exposure group.

c. Table 3 tabulates incidence and severity of lesions by organ observed in female rats dying spontaneously from the high (1000 mg/m³) dosage level exposure group.

III. DISCUSSION.

Numerous sporadically occurring lesions were noted in the high (1000 mg/m³) dosage level rats that died prior to completion of the White Phosphorus/Felt smoke inhalation studies. Moderate to severe laryngitis was observed in all of nine male rats examined and in five out of six female rats examined. Moderate to severe tracheitis was observed in five out of seven male rats and in all of nine female rats examined. Laryngeal and tracheal lesions often included varying degrees of vessiculation (blistering) and ulceration of the mucosal surfaces. Death of many of these animals probably resulted from asphyxiation produced by mechanical blockage of the larynx/trachea from swelling or from laryngospasm resulting from irritation of the larynx by the smoke. Moderate to severe congestion (hyperemia) was observed in twenty-five out of twenty-nine lungs examined. Irritation produced by the inhaled smoke may be responsible for this finding. A minimal to mild multifocal to diffuse interstitial pneumonia was observed in sixteen out of twenty-nine lungs examined. Peribronchiolar lymphoid aggregates were noted in the lungs of eighteen of twenty-nine animals examined. The interstitial pneumonia and peribronchiolar lymphoid aggregates have been observed historically in colony rats held in this and adjacent animal research facilities. The death of one control rat is attributed to a generalized Phycomycosis (non-contagious fungal agent) involving the gastrointestinal tract, liver and brain.

IV. CONCLUSION.

A high mortality rate was noted in the high (1000 mg/m³) dosage level exposure group of colony rats being exposed to White Phosphorus/Felt smoke via inhalation exposure. The death of these animals appears to be agent and dose related. A moderate to severe laryngitis/tracheitis accompanied by varying degrees of vessiculation (blistering) and ulceration was observed in most of the animals. Death in these animals probably resulted from asphyxiation produced by mechanical blockage of the larynx/trachea from tissue swelling or from laryngospasm resulting from irritation of the larynx by the smoke. Moderate to severe pulmonary congestion was observed in many of the animals and is probably agent related.



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Project Smoke II
White Phosphorus/Felt
Inhalation Study - Spontaneous Deaths
"Colony" Rats

Key to Microscopic Findings

Table 1

Number of Animals: Represents total number of animals in each group necropsied.

Numbers following tissues represent the number of tissues from each group actually examined.

Numbers following lesions indicate the number of tissues examined showing the particular lesion.

Tables 2-3

- = tissue present

+ = tissue not available for microscopic evaluation

Numbers following organ: Number of specimens examined

Severity of Response

1 = Minimal

2 = Slight

3 = Moderate

4 = Severe

P = Present

STRAIN OF RAT: COLONY
 EXPOSURE: VARIABLE
 AGENT: WHITE PHOSPHORUS/FELT

TABLE 1 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
 SMOKE II

NUMBER OF ANIMALS	CONTROL		HIGH DOSE	
	M	F	M	F
	0	1	15	14
NASAL TURBINATE	1/12/11		1	12/11
Rhinitis, Focal				2
LARYNX	1/9/6		1	9/6
Laryngitis				9/5
Submucosal Gland Duct Ectasia				
TRACHEA	0/7/9		7	9
Tracheitis				5/9
Submucosal Gland Duct Ectasia			1	
LUNGS	1/15/14		1	15/14
Pneumonia, Interstitial				8/8
Pneumonia, Granulomatous				
Pneumonia, Purulent			1	2
Congestion			12	13
Peribronchial Lymphoid Aggregates				9/9
Bronchitis				3

STRAIN OF RAT: COLONY
 EXPOSURE: VARIABLE
 AGENT: WHITE PHOSPHORUS/FELT

TABLE 1 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
 SMOKE II

NUMBER OF ANIMALS	CONTROL		HIGH DOSE	
	M	F	M	F
HEART	1/15/14		1	15/14
Myocardial Fibrosis, Focal				
Myocarditis, Focal			2	3
ESOPHAGUS	0/13/13	0	13	13
STOMACH	1/15/14		1	15/14
Gastritis, Acute				
SMALL INTESTINE	1/13/13		1	13/13
Enteritis, Acute, Ulcerative				
PANCREAS	1/14/14		1	14/14
LARGE INTESTINE	1/14/13		1	14/13
Hematochezia				
Enteritis, Acute			1	
LIVER	1/14/14		1	14/14
Hepatitis, Focal			1	
Infarct, Lobar				
ADRENAL	1/14/13		1	14/13

STRAIN OF RAT: COLONY
 EXPOSURE: VARIABLE
 AGENT: WHITE PHOSPHORUS/FELT

TABLE 1 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
 SMOKE II

NUMBER OF ANIMALS		CONTROL		HIGH DOSE	
		M	F	M	F
THYROID	0/10/13	0	0	10	13
Degeneration				8	11
THYMUS	0/13/7	0	0	13	7
Hemorrhage, Focal				1	
KIDNEY	1/15/14	1	1	15	14
Hydronephrosis					
Nephritis, Interstitial				2	
Glomerulonephritis				2	1
Tubular Mineralization, Focal				5	
Tubular Dilatation, Focal					
Proteinuria				4	5
BLADDER	0/9/14	0	0	9	14
Perivasculitis					
OVARY	0/0/14	0	0	0	4
UTERUS	1/0/11	1	1	0	1
Metritis					

STRAIN OF RAT: COLONY
 VARIABLE
 EXPOSURE: WHITE PHOSPHORUS/FELT
 AGENT:

TABLE 1 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM CONTROL AND SMOKE II

NUMBER OF ANIMALS	CONTROL		HIGH DOSE	
	M	F	M	F
MAMMARY GLAND	1/0/3		1	0 3
Adenoma				2
Mastitis				
TESTES	0/15/0		0	15 0
PROSTATE	0/11/0		0	11 0
Prostatitis				
MARROW	1/9/8		1	9 8
Granulopoiesis, Accelerated				
SPLEEN	1/15/14		1	15 14
Hemosiderosis				4
Lymphoid Deposition			1	
BRAIN	1/15/14		1	15 14
Encephalitis, Pyogranulomatous			1	
Necrosis, Focal				
EYE	1/13/13		1	13 13
PITUITARY	1/7/14		1	7 4
Cyst				1

STRAIN OF RAT: COLONY
EXPOSURE: VARIABLE
DOSE LEVEL: HIGH (1000 MG/M³)
AGENT: WHITE PHOSPHORUS/FELT

TABLE 2 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
SMOKE II

STRAIN OF RAT: COLONY
EXPOSURE: VARIABLE
DOSE LEVEL: HIGH (1000 MG/M³)
AGENT: WHITE PHOSPHORUS/FELT

TABLE 2 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
SMOKE II

STRAIN OF RAT: COLONY
EXPOSURE: VARIABLE
DOSE LEVEL: HIGH (1000 MG/M³)
AGENT: WHITE PHOSPHORUS/FELT

TABLE 2 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS SMOKE II

STRAIN OF RAT: COLONY
 EXPOSURE: VARIABLE
 DOSE LEVEL: HIGH (1000 MG/M³)
 AGENT: WHITE PHOSPHORUS/FELT

TABLE 2 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
 SMOKE II

	MALE														
	78-585	78-609	78-610	78-631	78-634	78-641	78-700	78-701	78-705	78-708	78-718	78-735	78-767	78-781	78-799
THYROID	10	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Degeneration		3	2	1			3	3	2				1	4	
THYMUS		13	-	+	+										
Hemorrhage, Focal															
KIDNEY		15	-	-	-	-	-	-	-	-	-				
Hydronephrosis															
Nephritis, Intersitial									3		1				
Glomerulonephritis															
Tubular Mineralization, Focal															
Tubular Dilatation, Focal															
Proteinuria															
BLADDER	9	-	-	+	-	-	+	+	-	+	-	-	-	-	-
Perivasculitis															
OVARY	3	+	+	+	+	+	+	+	+	+	+				
UTERUS	0	+	+	+	+	+	+	+	+	+	+				
Hemritis															3

STRAIN OF RAT: COLONY
EXPOSURE: VARIABLE
DOSE LEVEL: HIGH (1000 MG/M³)
AGENT: WHITE PHOSPHORUS/FELT

TABLE 3 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS SMOKE II

	NASAL TURBinate	Rhinitis, Focal	LARYNX	Laryngitis	Submucosal Gland Duct Ectasia	TRACHEA	Tracheitis.	Submucosal Gland Duct Ectasia	LUNGS	Pneumonia, Interstitial	Pneumonia, Granulomatous	Pneumonia, Purulent	Congestion	Peribronchial Lymphoid	Aggregates	Bronchitis
78-583	-	-	6	+	-	9	-	.	14	-	-	3	1	1	1	1
78-614A	-	-	+	+	-	+	+	-	-	2	2	2	2	2	2	2
78-620	-	-	4	4	4	3	4	3	1	1	3	3	3	3	3	3
78-621	-	-	4	4	4	3	4	4	1	1	4	4	4	4	4	4
78-622	-	-	4	4	4	3	4	4	1	1	4	4	4	4	4	4
78-642	-	-	4	4	4	3	4	4	1	1	4	4	4	4	4	4
78-643	-	-	4	4	4	3	4	4	1	1	4	4	4	4	4	4
78-646	-	-	4	4	4	3	4	4	1	1	4	4	4	4	4	4
78-665	-	-	4	4	4	3	4	4	1	1	4	4	4	4	4	4
78-703	-	-	4	4	4	3	4	4	1	1	4	4	4	4	4	4
78-709	-	-	4	4	4	3	4	4	1	1	4	4	4	4	4	4
78-724	-	-	4	4	4	3	4	4	1	1	4	4	4	4	4	4
78-730	-	-	4	4	4	3	4	4	1	1	4	4	4	4	4	4
78-737	-	-	4	4	4	3	4	4	1	1	4	4	4	4	4	4

STRAIN OF RAT: COLONY
 EXPOSURE: VARIABLE
 DOSE LEVEL: HIGH (1000 MG/M³)
 AGENT: WHITE PHOSPHORUS/FELT

TABLE 3 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
 SMOKE II

		FEMALE											
		78-583	78-614A	78-620	78-622	78-640	78-642	78-643	78-665	78-703	78-709	78-724	78-737
HEART	14	-	-	-	-	-	-	-	-	-	-	-	-
Myocardial Fibrosis, Focal													
Myocarditis, Focal	3			2		2							
ESOPHAGUS	13	-	-	-	+	-	-	-	-	-	-	-	-
Stomach	14	-	-	-	-	-	-	-	-	-	-	-	-
SMALL INTESTINE	13	-	-	-	-	-	-	-	-	-	-	-	-
PANCREAS	14	-	-	-	-	-	-	-	-	-	-	-	-
LARGE INTESTINE	13	-	-	-	-	-	-	-	-	-	-	-	-
Nematodiasis													
Enteritis, Acute													
LIVER	14	-	-	-	-	-	-	-	-	-	-	-	-
Hepatitis, Focal													
Infarct, Lobar													
ADRENAL	13	-	-	-	+	-	-	-	-	-	-	-	-

STRAIN OF RAT: COLONY
 EXPOSURE: VARIABLE
 DOSE LEVEL: HIGH (1000 MG/M³)
 AGENT: WHITE PHOSPHORUS/FELT

TABLE 3 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
 SMOKE II

		FEMALE					
THYROID	13	-	-	-	-	-	-
Degeneration		2	1	2	3	2	1
THYMUS	7	-	+	+	+	-	-
Hemorrhage, Focal							
KIDNEY	14	-	-	-	-	-	-
Hydronephrosis							
Nephritis, Interstitial		1	2				
Glomerulonephritis			2				
Tubular Mineralization, Focal		1	1		2	1	2
Tubular Dilatation, Focal							
Proteinuria							
BLADDER	14	-	-	-	-	-	-
Perivasculitis							
OVARY	14	-	-	-	-	-	-
UTERUS	11	-	-	-	-	-	+
Vaginitis							

STRAIN OF RAT: COLONY
EXPOSURE: VARIABLE
DOSE LEVEL: HIGH (7000 MG/M³)
AGENT: WHITE PHOSPHORUS/FELT

TABLE 3 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS SMOKE LI

APPENDIX D
PATHOLOGY REPORT - 6-WEEK EXPOSURE

Project Smoke II
Inhalation Study Six Weeks
"Colony" Rats

1. Introduction.

The present study was designed to assess potential local and toxic effects following inhalation exposure to White Phosphorus/Felt for a period of fifteen minutes per day, five days a week, for six weeks.

A group of six female rats were exposed by the inhalation route to White Phosphorus/Felt at a high dose level (1000 mg/m^3) and two groups of twelve rats, equally divided as to sex, received intermediate (500 mg/m^3) and low (200 mg/m^3) dosage levels for the prescribed period of time followed by euthanasia. A fourth group of eighteen rats, equally divided as to sex, that were exposed to air only and maintained under similar conditions, served as controls.

At termination and necropsy, tissues were imbedded in paraffin and subsequently processed for staining with hematoxylin and eosin. The following tissues were evaluated microscopically: nasal turbinate, larynx, trachea, lungs, heart, esophagus, stomach, small intestine, pancreas, large intestine, liver, adrenal, thyroid, thymus, kidney, bladder, ovary/teste, uterus, mammary gland, prostate, bone marrow, spleen, brain, eye and pituitary.

2. Results.

The microscopic observations are presented in the Histopathology Incidence Tables.

a. Tables 1-4 tabulate incidence of lesions (by organ) observed in the four groups of male and female colony rats.

- b. Tables 5-8 tabulate incidence and severity of lesions (by organ) observed in male and female control rats.
- c. Tables 9-12 tabulate incidence of lesions (by organ) observed in each female rat receiving the high dose level exposure to White Phosphorus/Felt.
- d. Tables 13-16 tabulate incidence of lesions (by organ) observed in each male and female rat receiving the intermediate dose level exposure to White Phosphorus/Felt.

3. Discussion.

A number of spontaneous lesions were noted in the liver, kidney, thyroid, and pituitary; the lesions occurring with equal severity and frequency in control and White Phosphorus/Felt exposed rats. While none of the control animals displayed laryngitis or tracheitis, all of the laryngeal and tracheal specimens examined from the rats receiving the high dosage level of White Phosphorus/Felt displayed a moderate to severe laryngitis/tracheitis. Fifty percent of the rats receiving intermediate dose levels of White Phosphorus/Felt displayed a minimal to mild tracheitis while one out of three rats had a mild laryngitis. Only one rat receiving the low dose level of White Phosphorus/Felt displayed tracheitis. Four out of six female rats receiving the high dose level of White Phosphorus/Felt displayed minimal to severe interstitial pneumonia while one control rat of each sex displayed a minimal interstitial pneumonia.

4. Conclusion.

The agent, White Phosphorus/Felt, at the dosage levels tested and in the manner tested show a dose related laryngitis and tracheitis. It is suggestive

that high dosage levels of White Phosphorus/Felt may produce interstitial pneumonia in some animals.

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4 September 1979

Project Smoke II
White Phosphorus Felt
Inhalation Study Six Weeks
"Colony" Rats

Key to Microscopic Findings

Tables 1-4

Number of Animals: Represents total number of animals in each group utilized in the study.

Numbers following tissues represent the number of tissues from each group actually examined.

Numbers following lesion indicate the number of tissues examined showing the particular lesion.

Tables 5-16

- = tissue present
* = tissue not available for microscopic evaluation

Severity of Response

1 = minimal
2 = slight
3 = moderate
4 = severe
P = present

TABLE I: MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
SMOKE II, 6 WEEKS EXPOSURE

STRAIN OF RAT: COLONY
WEEKS EXPOSURE: 6
DAYS POST EXPOSURE: 1
AGENT: WHITE PHOSPHORUS FELT

NUMBER OF ANIMALS	CONTROL			HIGH			INTERMEDIATE			LOW		
	M	F	G	M	F	G	M	F	G	M	F	G
NASAL TURBinate	7	8		5	4		4	3		6	6	
Rhinitis, Focal												
LARYNX	7	4		2	0		3	0		2		
Laryngitis										1		
Submucosal Gland Duct Ectasia	3	2		1								
TRACHEA	3	7		4	6		4	6		6	3	
Tracheitis												
Submucosal Gland Duct Ectasia	2			4	3		2	1		3	1	
LUNGS	9	9		6	6		6	6		6	6	
Pneumonia, Interstitial	1	1		4						2		
Pneumonia, Granulomatous												
Pneumonia, Purulent										2		
Congestion							5	5		3	5	
Peribronchiolar Lymphoid	6	4										
Aggregates												
Bronchitis												

STRAIN OF RAT: COLONY
 WEEKS EXPOSURE: 6
 DAYS POST EXPOSURE: 1
 AGENT: WHITE PHOSPHORUS FELT

TABLE 2: MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS

NUMBER OF ANIMALS	CONTROL		HIGH		INTERMEDIATE		LOW	
	M	F	M	F	M	F	M	F
HEART	9	9	0	6	6	6	6	6
Myocardial Fibrosis, Focal	7	1	6	6	6	6	6	6
Myocarditis, Focal	3							
ESOPHAGUS	9	8	5	5	6	6	6	6
STOMACH	9	9	6	6	5	5	5	6
SMALL INTESTINE	9	9	5	6	6	6	6	4
PANCREAS	9	9	6	5	5	5	6	6
LARGE INTESTINE	9	9	6	4	6	6	6	6
Nematodiasis	1				1			
Enteritis, Acute								
LIVER	9	9	6	6	6	6	6	6
Hepatitis, Focal	3							
Infarct, Lobar								
ADRENALE	9	9	6	5	5	5	6	6

TABLE 3: MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
STRAIN OF RAT: COLONY
WEEKS EXPOSURE: 6
DAYS POST EXPOSURE: 1
AGENT: WHITE PHOSPHORUS FELT

NUMBER OF ANIMALS	CONTROL			HIGH			INTERMEDIATE			LOW		
	M	F	M	M	F	M	M	F	M	F	M	F
THYROID	8	7			4		5	3			5	6
Degeneration	7	4		2		3	2		2		2	3
THYMUS	9	7		4		5	5		5		5	5
Hemorrhage, Focal												
KIDNEY	9	9		6		6	6		6		6	6
Hydronephrosis	1	1										
Nephritis, Interstitial			1		1							
Glomerulonephritis				6	5		5	1	1			
Tubular Mineralization, Focal				1								
Tubular Dilatation, Focal				1		1		1				
Proteinuria				1	1							
BLADDER	7	7			3							
Perivasculitis												
OVARY	0	9		6		5		5	0	6		
UTERUS			8		6		4	0	4			
Hemritis								1				

TABLE 4: MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
SMOKE II, 6 WEEKS EXPOSURE:

STRAIN OF RAT: COLONY
WEEKS EXPOSURE: 6
DAYS POST EXPOSURE: 1
AGENT: WHITE PHOSPHORUS FELT

	NUMBER OF ANIMALS		CONTROL		HIGH		INTERMEDIATE		LOW	
	M	F	M	F	M	F	M	F	M	F
MAMMARY GLAND	0	2		0	6	6	6	6	6	6
Adenoma									0	1
Mastitis										
TESTES	9	0	0	0	6	0	6	0	6	0
PROSTATE	9	0	0	0	4	0	4	0	4	0
Prostatitis										
MARROW	6	8	5	5	4	4	6	6	6	6
Granulopoeisis, Accelerated	1		1		2					
SPLEEN	9	8	6	6	6	6	6	6	6	6
Hemosiderosis									3	1
BRAIN	9	9	6	5	5	6	6	6	6	6
Encephalitis, Pyogranulomatous										
Necrosis, Focal	7	9	5	5	5	5	5	5	5	6
EYE									4	4
PITUITARY	5	7	6	6	4	4	4	4	6	6
Adenoma		1								4

STRAIN OF RAT: COLONY
 WEEKS EXPOSURE: 6
 DAYS POST EXPOSURE: 1
 DOSE LEVEL: Control
 NECROPSY NO: 78-800/801/731/730/745/746
 AGENT: WHITE PHOSPHORUS FELT

Appendix D

TABLE 5: MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
SMOKE II, 6 WEEKS EXPOSURE

	MALE										FEMALE									
	A	B	C	D	E	F	G	H	I	J	A	B	C	D	E	F	G	H	I	
NASAL TURBinate	-	*	-	-	*	-	-	-	-	-	-	*	-	-	-	-	*	-	-	-
Rhinitis, Focal	-	-	-	-	-	-	-	-	-	-	-	*	*	*	*	*	-	-	*	-
LARYNX	*	-	*	-	-	-	-	-	-	-	*	*	*	*	*	*	*	*	*	*
Laryngitis	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Submucosal Gland Duct Ectasia	-	-	-	-	-	-	-	-	-	-	1	3	1	3	2	-	-	-	-	-
TRACHEA	-	-	*	*	*	*	*	*	*	*	-	*	*	*	*	*	*	*	*	*
Tracheitis	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Submucosal Gland Duct Ectasia	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
LUNGS	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Pneumonia, Interstitial	-	-	-	-	-	-	-	-	-	-	2	-	-	-	-	-	-	-	-	-
Pneumonia, Granulomatous	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Pneumonia, Purulent	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Congestion	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Peribronchiolar Lymphoid	-	-	-	-	-	-	-	-	-	-	1	1	1	1	1	1	1	1	1	1
Aggregates	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Bronchitis	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

STRAIN OF RAT: COLONY
 WEEKS EXPOSURE: 6
 DAYS POST EXPOSURE: 1
 DOSE LEVEL: Control
 NECROPSY NO: 78-800/801/731/730/745/746
 AGENT: WHITE PHOSPHORUS FELT

TABLE 6: MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
SMOKE II, 6 WEEKS EXPOSURE

	MALE										FEMALE									
	A	B	C	D	E	F	G	H	I	J	A	B	C	D	E	F	G	H	I	J
HEART	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Myocardial Fibrosis, Focal																				
Myocarditis, Focal	2										1	1								
ESOPHAGUS	-	-	-	-	-	-	-	-	-	-	-	*	-	*	-	-	-	-	-	-
STOMACH	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
SMALL INTESTINE	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
PANCREAS	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
LARGE INTESTINE	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Nematodiasis															P					
Enteritis, Acute																				
LIVER	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Hepatitis, Focal																		1	1	1
Infarct, Lobar																				
ADRENAL	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

2

STRAIN OF RAT: COLONY
 WEEKS EXPOSURE: 6
 DAYS POST EXPOSURE: 1
 DOSE LEVEL: Control
 NECROPSY NO: 78-801/731/730/745/746
 AGENT: WHITE PHOSPHORUS FELT

TABLE 7: MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
SMOKE II, 6 WEEKS EXPOSURE

	MALE										FEMALE										
	A	B	C	D	E	F	G	H	I	J	A	B	C	D	E	F	G	H	I	J	
THYROID	*	-	-	-	-	-	-	-	*	-	-	*	-	-	*	-	-	-	-	-	
Degeneration		4	3	3	1	2	2	2				4	2	4	2	4	3				
THYMUS									*	*											
Hemorrhage, Focal																					
KIDNEY																					
Hydronephrosis																					
Nephritis, Interstitial																					
Glomerulonephritis																					
Tubular Mineralization, Focal												2	1	1	1	1	1	1	1	1	2
Tubular Dilatation, Focal												1	1	1	1	1	1	1	1	1	1
Proteinuria																					
BLADDER									*	-											
Perivasculitis																					
OVARY																					
UTERUS																					3
Metritis																					

STRAIN OF RAT: COLONY
 WEEKS EXPOSURE: 6
 DAYS POST EXPOSURE: 1
 DOSE LEVEL: Control
 NECROPSY NO: 78-800/801/731/730/745/746
 AGENT: WHITE PHOSPHORUS FELT

TABLE 8: MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS

	FEMALE										MALE									
	A	B	C	D	E	F	G	H	I	J	A	B	C	D	E	F	G	H	I	J
MAMMARY GLAND											*	*	*	*	*	*	*	*	*	*
Adenoma											-	-	-	-	-	-	-	-	-	-
Mastitis											-	-	-	-	-	-	-	-	-	-
TESTES											-	-	-	-	-	-	-	-	-	-
PROSTATE											-	-	-	-	-	-	-	-	-	-
Prostatitis											*	*	*	*	*	*	*	*	*	*
MARROW		*	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Granulopoiesis, Accelerated											2	-	-	-	-	-	-	-	-	-
SPLEEN											*	-	-	-	-	-	-	-	-	-
Hemosiderosis											-	2	2	1	2	1	2	1	2	1
BRAIN											-	-	-	-	-	-	-	-	-	-
Encephalitis, Pyogranulomatous											-	-	-	-	-	-	-	-	-	-
Necrosis, Focal	*	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
EYE	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*
PITUITARY	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*
Adenoma											-	-	-	-	-	-	-	-	-	-

STRAIN OF RAT: COLONY
 WEEKS EXPOSURE: 6
 DAYS POST EXPOSURE: 1
 DOSE LEVEL: HIGH
 RECROPSY NO: 78-728/729
 AGENT: WHITE PHOSPHORUS FELT

Appendix D

TABLE 9: MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
SMOKE II, 6 WEEKS EXPOSURE

	MALE										FEMALE									
	A	B	C	D	E	F	G	H	I	J	A	B	C	D	E	F	G	H	I	J
NASAL TURBinate	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Rhinitis, Focal	-	*	*	*	*	*	*	*	*	*	-	*	*	*	*	*	*	*	*	*
LARYNX											3									3
Laryngitis											3									
Submucosal Gland Duct Ectasia											*	*								
TRACHEA											3	3	4	4						
Tracheitis.																				
Submucosal Gland Duct Ectasia																				
LUNGS											-	-	-	-	-	-	-	-	-	-
Pneumonia, Interstitial											3	4	3	1						
Pneumonia, Granulomatous																				2
Pneumonia, Purulent																				3
Congestion																				2
Peribronchiolar Lymphoid																				2
Aggregates																				2
Bronchitis																				1

STRAIN OF RAT: COLONY
 WEEKS EXPOSURE: 6
 DAYS POST EXPOSURE: 1
 DOSE LEVEL: High
 NECROPSY NO: 78-728/729
 AGENT: WHITE PHOSPHORUS FELT

TABLE 10: MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS

SMOKE II, 6 WEEKS EXPOSURE

MALE

FEMALE

	A	B	C	D	E	F	G	H	I	J	A	B	C	D	E	F	G	H	I	J
HEART	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Myocardial Fibrosis, Focal	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*
Myocarditis, Focal	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*
ESOPHAGUS	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*
STOMACH	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
SMALL INTESTINE	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*
PANCREAS	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
LARGE INTESTINE	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
Nematodiasis	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
Enteritis, Acute	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
LIVER	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
Hepatitis, Focal	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
Infarct, Lobar	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
ADRENAL	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	

STRAIN OF RAT: COLONY
 WEEKS EXPOSURE: 6
 DAYS POST EXPOSURE: 7
 DOSE LEVEL: High
 NECROPSY NO: 78-728/729
 AGENT: WHITE PHOSPHORUS FELT

TABLE II: MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS

	MALE										FEMALE									
	A	B	C	D	E	F	G	H	I	J	A	B	C	D	E	F	G	H	I	J
THYROID	-	-	-	-	-	-	-	-	-	*	-	-	-	-	-	-	-	-	-	-
Degeneration	-	-	-	-	-	-	-	-	-	*	-	-	-	-	-	-	-	-	-	-
THYMUS	-	-	-	-	-	-	-	-	-	*	-	-	-	-	-	-	-	-	-	*
Hemorrhage, Focal	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
KIDNEY	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Hydronephrosis	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Nephritis, Interstitial	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Glomerulonephritis	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Tubular Mineralization, Focal	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Tubular Dilatation, Focal	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Proteinuria	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
BLADDER	-	-	-	-	-	-	-	-	-	*	-	*	-	*	-	*	-	*	-	-
Perivasculitis	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
OVARY	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
UTERUS	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Hysteritis	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	3

STRAIN OF RAT: COLONY 6
 WEEKS EXPOSURE: 6
 DAYS POST EXPOSURE: 1
 DOSE LEVEL: High
 NECROPSY NO: 78-728/729
 AGENT: WHITE PHOSPHORUS FELT

TABLE 12: MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS

	MALE										FEMALE									
	A	B	C	D	E	F	G	H	I	J	A	B	C	D	E	F	G	H	I	J
MAMMARY GLAND											*	-	*	-	*	-	*	-	*	-
Adenoma																				
Mastitis																				
TESTES											*	*	*	*	*	*	*	*	*	*
PROSTATE											*	*	*	*	*	*	*	*	*	*
Prostatitis																				
MARROW											-	-	*	-	-	-	-	-	-	-
Granulopoiesis, Accelerated																	2			
SPLEEN											-	-	-	-	-	-	-	-	-	-
Hemosiderosis																		1	1	
BRAIN																				
Encephalitis, Pyogranulomatous																				
Necrosis, Focal																				
EYE																				
PITUITARY																				
Adenoma																				

STRAIN OF RAT: COLONY
 WEEKS POST EXPOSURE: 6
 DAYS POST EXPOSURE: 1
 DOSE LEVEL: Intermediate
 NECROPSY NO: 78-747/748
 AGENT: WHITE PHOSPHORUS FELT

Appendix D

TABLE 13: MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS

	MALE										FEMALE										
	A	B	C	D	E	F	G	H	I	J	A	B	C	D	E	F	G	H	I	J	
NASAL TURBinate	-	*	-	*	-	-	*	-	*	-	-	*	*	-	*	*	-	*	-	*	
Rhinitis, Focal																					
LARYNX	*	*	*	*	*	*	*	*	*	*	-	*	-	*	*	*	-	*	*	*	*
Laryngitis																					2
Submucosal Gland Duct Ectasia																					3
TRACHEA	-	-	-	-	-	-	-	-	-	-	-	*	-	*	-	*	-	*	-	-	
Tracheitis																					1
Submucosal Gland Duct Ectasia																					
LUNGS	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
Pneumonia, Interstitial																					4
Pneumonia, Granulationous																					
Pneumonia, Purulent																					
Congestion																					
Peribronchiolar Lymphoid	1	1	1	2	1																1
Aggregates																					
Bronchitis																					

STRAIN OF RAT: COLONY
 WEEKS EXPOSURE: 6
 DAYS POST EXPOSURE: 1
 DOSE LEVEL: Intermediate
 NECROPSY NO: 78-747/742
 AGENT: WHITE PHOSPHORUS FELT

TABLE 14: MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS

	MALE										FEMALE										
	A	B	C	D	E	F	G	H	I	J	A	B	C	D	E	F	G	H	I	J	
HEART	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
Myocardial Fibrosis, Focal											2										
Myocarditis, Focal											-										
ESOPHAGUS											-										
STOMACH											-										
SHELL INTESTINE											-										
PANCREAS											-	*									
LARGE INTESTINE											-	-									
Kryptosporidiosis											P										
Enteritis, Acute											-	-	-	-	-	-	-	-	-	-	
LIVER											-	-	-	-	-	-	-	-	-	-	
Hepatitis, Focal											-										
Infiltrat, Lobar											-	*									
ADRENAL											-	*									

STRAIN OF RAT: COLONY
 WEEKS EXPOSURE: 6
 DAYS POST EXPOSURE: 1
 DOSE LEVEL: Intermediate
 NECROPSY NO: 78-747748
 AGENT: WHITE PHOSPHORUS FELT

TABLE 15: MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
SMOKE II, 6 WEEKS EXPOSURE

	MALE										FEMALE										
	A	B	C	D	E	F	G	H	I	J	A	B	C	D	E	F	G	H	I	J	
THYROID	-	-	*	-	-	-	-	*	*	*	-	*	*	*	*	-	-	-	-	-	
Degeneration						4	3	2								4	3				
THYMUS	*	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	*	-	
Hemorrhage, Focal						1															
KIDNEY	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
Hydronephrosis																					
Nephritis, Interstitial																					
Glomerulonephritis																					
Tubular Mineralization, Focal																	2	1	2	1	1
Tubular Dilatation, Focal																					
Proteinuria											1										
BLADDER	*	*	*	*	-	-	-	-	-	-	-	*	-	*	-	*	-	-	*	-	-
Perivasculitis																					
OVARY																					
UTERUS																					
Metritis																					

STRAIN OF RAT: COLONY
 WEEKS EXPOSURE: 6
 DAYS POST EXPOSURE: 1
 DOSE LEVEL: Intermediate
 NECROPSY NO: 78-747/743
 AGENT: WHITE PHOSPHORUS FELT

TABLE 16: MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS

	MALE										FEMALE									
	A	B	C	D	E	F	G	H	I	J	A	B	C	D	E	F	G	H	I	J
MAMMARY GLAND											*	-	-	*	*	*	*	*	*	*
Adenoma																				
Mastitis																				
TESTES											-									
PROSTATE											-	*	-	*						
Prostatitis																				
MARROW											-	*	-	*						
Granulopoiesis, Accelerated																	3	2		
SPLEEN											-	-	-	-	-	-	-	-	-	
Hemosiderosis												-	-	-	-	-	-	-	-	
BRAIN																				
Encephalitis, Pyogranulomatous																				
Necrosis, Focal																				
EYE																				
PITUITARY																				
Adenoma																				

STRAIN OF RAT: COLONY
 WEEKS EXPOSURE: 6
 DAYS POST EXPOSURE: 1
 DOSE LEVEL: LOW
 NECROPSY NO: 78-796/798
 AGENT: WHITE PHOSPHORUS FELT

Appendix D

TABLE 17: MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
SMOKE II, 6 WEEKS EXPOSURE

	FEMALE										MALE											
	A	B	C	D	E	F	G	H	I	J	A	B	C	D	E	F	G	H	I	J		
NASAL TURBinate	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-		
Rhinitis, Focal																						
LARYNX	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	*	
Laryngitis																						
Submucosal Gland Duct Ectasia																						
TRACHEA	-	-	-	-	-	-	-	-	-	-	-	*	*	*	*	*	*	*	*	-	-	
Tracheitis																						
Submucosal Gland Duct Ectasia																						
LUNGS	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
Pneumonia, Interstitial																						
Pneumonia, Granulomatous																						
Pneumonia, Purulent																						
Congestion																						
Peribronchial Lymphoid Aggregates	1	1	2									1	1	1	1	1	1	1	1	1	1	
Bronchitis																						

STRAIN OF RAT: COLONY
 WEEKS EXPOSURE: 6
 DAYS POST EXPOSURE: 1
 DOSE LEVEL: LOW
 NECROPSY NO: 78-796/798
 AGENT: WHITE PHOSPHORUS - FELT

TABLE 18: MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS

	MALE										FEMALE									
	A	B	C	D	E	F	G	H	I	J	A	B	C	D	E	F	G	H	I	J
HEART	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Myocardial Fibrosis, Focal																				
Myocarditis, Focal	1																			
ESOPHAGUS	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
STOMACH	-	-	*	-	-	-	-	-	-	-	-	-	-	*	-	-	-	-	-	-
SMALL INTESTINE	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
PANCREAS	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
LARGE INTESTINE	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Nematodiasis																				
Enteritis, Acute																				
LIVER	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Hepatitis, Focal																				
Infarct, Lobar																				
ADRENAL	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	2

STRAIN OF RAT: COLONY
 WEEKS EXPOSURE: 6
 DAYS POST EXPOSURE: 1
 DOSE LEVEL: LOW
 NECROPSY NO: 78-796/798
 AGENT: WHITE PHOSPHORUS FELT

TABLE 19: MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
SMOKE II, 6 WEEKS EXPOSURE

	MALE										FEMALE										
	A	B	C	D	E	F	G	H	I	J	A	B	C	D	E	F	G	H	I	J	
THYROID	-	-	*	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
Degeneration			3	2							3	1	2								
THYMUS	*	-	-	-	-	-	-	-	-	-	-	-	*	-	-	-	-	-	-	-	
Hemorrhage, Focal																					
KIDNEY																					
Hydronephrosis																					
Nephritis, Interstitial																					
Glomerulonephritis																					
Tubular Mineralization, Focal																					
Tubular Dilatation, Focal																					
Proteinuria																					
BLADDER	-		*	*	-	*	*														
Perivasculitis																					
OVARY																					
UTERUS																					
Metritis																					3

STRAIN OF RAT: COLONY
 WEEKS EXPOSURE: 6
 DAYS POST EXPOSURE: 1
 DOSE LEVEL: Low
 NECROPSY NO: 78-796/798
 AGENT: WHITE PHOSPHORUS FELT

TABLE 20: MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS

SMOKE 11, 6 WEEKS EXPOSURE

	MALE										FEMALE									
	A	B	C	D	E	F	G	H	I	J	A	B	C	D	E	F	G	H	I	J
MAMMARY GLAND											*	*	-	*	*	*	*	*	*	*
Adenoma																				
Mastitis																				
TESTES																				
PROSTATE																				
Prostatitis																				
MARROW																				
Granulopoiesis, Accelerated																				
SPLEEN																				
Hemosiderosis																				
BRAIN																				
Encephalitis, Pyogranulomatous																				
Necrosis, Focal																				
EYE																				
PITUITARY																				
Adenoma																				

APPENDIX E
PATHOLOGY REPORT - 13-WEEK EXPOSURE

PATHOLOGY REPORT
PROJECT SMOKE II, WHITE PHOSPHORUS/FELT
INHALATION STUDY - THIRTEEN WEEKS
"COLONY" RATS

I. INTRODUCTION.

The study was designed to assess potential local and systemic toxic effects on Edgewood Area Colony rats following inhalation exposure in a chamber to White Phosphorus/Felt for a period of fifteen minutes per day, five days a week, for thirteen weeks. Three separate dosage levels were studied with each dosage level beginning on a different date and having a separate set of control animals receiving sham treatment and housing under similar conditions. All animals were approximately six weeks of age at the beginning of each study. The exposures and animal holding were performed in Bldg E3266.

Exposure of the high dose (1000 mg/m^3) group began on 31 July and ended on 31 October 1978. The exposed group consisted of eight animals, four of each sex. A separate group of three male and three female controls were utilized. Exposure of the medium dose (500 mg/m^3) group began on 7 August 1978 and ended on 7 November of that year. Twelve rats, six of each sex, were exposed to the agent. Three male and three female control rats were utilized in this study. Exposure of the low dose (200 mg/m^3) began on 11 September 1978 and ended on 12 December of the same year. A group of twelve animals, six of each sex, were exposed to the agent. Three male and three female controls were utilized in the study.

In addition to the above animals, a total of four rats, two of each sex, that had been exposed with the high dose group and utilized in physiology studies for eight days were necropsied. No controls were submitted with this group of animals. These animals were included separately in Tables 17-20, but are not considered part of the study in interpreting pathology findings.

Following necropsy, tissues were imbedded in paraffin and subsequently processed for staining with hematoxylin and eosin. The following tissues were evaluated microscopically: nasal turbinates, larynx, trachea, lungs, heart, esophagus, stomach, small intestine, pancreas, large intestine, liver, adrenal, thyroid, thymus, kidney, bladder, ovary/teste, uterus, mammary gland, prostate, bone marrow, spleen, brain, eye and pituitary.

Histologic findings are tabulated in Tables 1-20. Since one or more tissues from various animals were lost at necropsy or during processing, one must calculate the incidence of lesions based upon the number of tissues examined rather than on the number of animals necropsied.

II. RESULTS.

The microscopic observations are presented in the Histopathology Incidence Tables.

a. Tables 1-4 tabulate incidence of lesions (by organ) observed in the six groups of male and female colony rats utilized in the three studies.

b. Tables 5-8 tabulate incidence and severity of lesions (by organ) observed in male and female control rats.

c. Tables 9-12 tabulate incidence of lesions (by organ) observed in each male and female rat receiving the high dose (1000 mg/m^3) level exposure to White Phosphorus/Felt.

d. Tables 13-16 tabulate incidence of lesions (by organ) observed in each male and female rat receiving the medium dose (500 mg/m^3) level exposure.

e. Tables 17-20 tabulate incidence of lesions (by organ) observed in each male and female rat receiving the low dose (200 mg/m^3) level exposure.

III. DISCUSSION.

A number of spontaneous lesions were noted in the liver, kidney, thyroid, spleen, heart, uterus, bladder, mammary glands and large intestine. The lesions occurred either sporadically or with equal frequency and severity in both control and White Phosphorus/Felt exposed rats. While none of the control rats displayed laryngitis or tracheitis, all of the male rats receiving the high dosage level of White Phosphorus/Felt exhibited a moderate laryngitis. Of the female rats receiving the same dosage level (1000 mg/m^3) of White Phosphorus/Felt, one of two larynges examined displayed a moderate laryngitis while two out of three tracheae displayed mild to moderate tracheitis. Three out of six female rats receiving the medium dosage level of White Phosphorus/Felt displayed a moderate tracheitis while three out of five male rats receiving the same level exposure displayed slight to moderate tracheitis. None of the low dosage level animals displayed laryngitis or tracheitis.

IV. CONCLUSION.

Though the small number of tissues examined prevents a definitive conclusion, it appears that the agent, White Phosphorus/Felt produces a dose related laryngitis and tracheitis when rats are exposed in the manner tested.



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Project Smoke II
White Phosphorus/Felt
Inhalation Study - Thirteen Weeks
"Colony" Rats

Key to Microscopic Findings

Tables 1-4

Number of Animals: Represents total number of animals in each group utilized in the study.

Numbers following tissues represent the number of tissues from each group actually examined.

Numbers following lesions indicate the number of tissues examined showing the particular lesion.

Tables 5-20

- = tissue present

+ = tissue not available for microscopic evaluation

Numbers Following Organs: First number is number of tissues from males examined, second number is number of tissues from females.

Severity of Response

1 = Minimal

2 = Slight

3 = Moderate

4 = Severe

P = Present

STRAIN OF RAT: Colony
 EXPOSURE: 13 wks
 AGENT: White Phosphorus/Feit

TABLE I : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
 SMOKE II

NUMBER OF ANIMALS	HIGH DOSE						MEDIUM DOSE						LOW DOSE					
	CONTROL			EXPOSED			CONTROL			EXPOSED			CONTROL			EXPOSED		
	M	F	M	F	M	F	M	F	M	F	M	F	M	F	M	F	M	F
NASAL TURBINATE	3	3	4	4	3	3	6	6	3	3	6	6	3	3	6	6	3	3
Rhinitis, Focal	3	2	3	3	3	2	4	6	2	3	4	6	2	3	5	6	2	3
LARYNX	3	2	2	2	0	1	0	0	2	0	0	0	1	4	1	4	1	4
Laryngitis			2	1										4				
Submucosal Gland Duct Ectasia	1	1																
TRACHEA	1	2	0	3	3	3	5	6	3	3	5	6	3	3	5	4	3	4
Tracheitis			2						3	3			1					
Submucosal Gland Duct Ectasia																		
LUNGS	3	3	4	4	3	3	6	6	3	3	6	6	3	3	6	6	3	3
Pneumonia, Interstitial		1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1
Pneumonia, Granulomatous																		
Pneumonia, Purulent																		
Congestion	3	1	2		3	2	5	6	3	2	4	5	3	2	4	5	3	2
Peribronchiolar Lymphoid Aggregates																		
Bronchitis																		

STRAIN OF RAT: Colony
 EXPOSURE: 13 wks
 AGENT: White Phosphorus/Felt

TABLE 2 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
 SMOKE II

NUMBER OF ANIMALS	HIGH DOSE						MEDIUM DOSE						LOW DOSE					
	CONTROL			EXPOSED			CONTROL			EXPOSED			CONTROL			EXPOSED		
	M	F	M	F	M	F	M	F	M	F	M	F	M	F	M	F	H	F
HEART	3	3	4	4	3	3	6	6	3	3	3	3	6	6	3	3	6	6
Myocardial Fibrosis, Focal	3	3	4	4	3	3	6	6	3	3	3	3	6	6	3	3	6	6
Myocarditis, Focal			1				3		2									
ESOPHAGUS	3	3	3	3	3	3	4	6	3	3	5	4						
STOMACH	2	3	4	4	3	2	6	6	3	3	3	3	6	6				
SMALL INTESTINE	3	3	3	4	3	3	5	4	2	3	3	3	6	6				
PANCREAS	3	3	4	4	3	2	5	6	3	3	3	3	6	6				
LARGE INTESTINE	1	3	3	1	3	3	6	6	2	3	2	3	6	6				
Nematodiasis							1											
Enteritis, Acute																		
LIVER	3	3	4	4	3	3	5	6	3	3	3	3	6	6				
Hepatitis, Focal	1				1	1			1									
Infarct, Lobar																		
ADRENAL	3	3	3	4	2	2	6	6	3	3	3	3	6	6				

STRAIN OF RAT: Colony
EXPOSURE: 13 wks
AGENT: White Phosphorus/Fett

TABLE 3 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
SMOKE II

NUMBER OF ANIMALS	HIGH DOSE						MEDIUM DOSE						LOW DOSE					
	CONTROL			EXPOSED			CONTROL			EXPOSED			CONTROL			EXPOSED		
	M	N	F	M	N	F	M	N	F	M	N	F	M	N	F	M	N	F
THYROID	3	3	4	4	4	4	3	3	3	6	6	6	3	3	3	6	6	6
Degeneration	3	3	3	2	3	2	3	2	3	5	6	5	3	2	3	4	4	4
THYMUS	2	3	4	3	3	3	3	3	3	5	5	5	3	3	3	6	6	6
Hemorrhage, Focal																		
KIDNEY	3	3	4	4	4	4	3	3	3	6	6	6	3	3	3	6	6	6
Hydronephrosis																		
Nephritis, Interstitial	1						1						1					
Glomerulonephritis																		
Tubular Mineralization, Focal	1			1			1			2			3			3		
Tubular Dilatation, Focal										1								
Proteinuria	1									2						1		
BLADDER	1	1	4	4	0	1				5	2		3	1		5	4	
Perivasculitis																		
OVARY	0	2	0	3	0	3	0	3	0	5	0	3	0	3	0	6	0	6
UTERUS	0	2	0	4	0	2	0	2	0	4	0	4	0	3	0	5	0	5
Hepatitis																1		

STRAIN OF RAT: Colony
 EXPOSURE: 12 wks
 AGENT: White Phosphorus/F₃C

TABLE 4 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
 SMOKE II

NUMBER OF ANIMALS	HIGH DOSE			MEDIUM DOSE			LOW DOSE			
	CONTROL		EXPOSED	CONTROL		EXPOSED	CONTROL		EXPOSED	
	M	F	H	F	H	F	H	F	H	F
MAMMARY GLAND	3	0	4	4	3	3	5	6	3	6
Adenoma	0	3	3	2	0	3	0	3	0	3
Mastitis	3	0	4	0	3	0	6	0	3	0
TESTES	0	0	4	0	3	0	6	0	3	0
PROSTATE	0	0	4	0	3	0	6	0	3	0
Prostatitis									2	
MARROW	3	2	4	3	3	3	5	6	3	3
Granulopoiesis, Accelerated									1	
SPLEEN	2	2	4	4	3	3	6	5	2	3
Hemosiderosis	2	2	3	3			3	4	3	4
BRAIN	3	3	4	4	3	2	6	6	3	3
Encephalitis, Pyogranulomatous										
Necrosis, Focal	3	3	4	4	3	3	5	5	3	3
Eye	2	2	3	2	3	2	1	6	3	3
PITUITARY										
Adenoma										

STRAIN OF RAT: Colony
 EXPOSURE: 12 wks
 DOSE LEVEL: Control
 AGENT: White Phosphorus/Fest

TABLE 5 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
SMOKE II

	MALE			FEMALE		
	HIGH 78-833	MED 78-870	LOW 78-959	HIGH 78-834	MED 78-872	LOW 78-950
	A B C	A B C	A B C	A B C	A B C	A B C
NASAL TURBinate	-	-	-	-	-	-
Rhinitis, Focal						
LARYNX	-	-	-	-	-	-
Laryngitis						
Submucosal Gland Duct Ectasia		2		2		
TRACHEA	+	-	-	-	-	-
Tracheitis						
Submucosal Gland Duct Ectasia						
LUNGS	-	-	-	-	-	-
Pneumonia, Interstitial				2	2	2
Pneumonia, Granulomatous						
Pneumonia, Purulent						
Congestion						
Peribronchiolar Lymphoid	1	1	2	1	1	2
Aggregates						
Bronchitis						

STRAIN OF RAT: CO₂ only
 EXPOSURE: 13 wks.
 DOSE LEVEL: Control
 AGENT: White Phosphorus, f.e.t.

TABLE 6 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM DOLORY RATS

	MALE			FEMALE			LOW		
	HIGH	MED	LOW	HIGH	MED	LOW	HIGH	MED	LOW
	78-833	78-870	78-955	78-834	78-872	78-960			
	A	B	C	A	B	C	A	B	C
	A	B	C	A	B	C	A	B	C
<u>HEART</u>	-	-	-	-	-	-	-	-	-
<u>Myocardial Fibrosis, Focal</u>				+	+	+			
<u>Myocarditis, Focal</u>				+	+	+			
<u>ESOPHAGUS</u>	-	-	-	-	-	-	-	-	-
<u>STOMACH</u>	-	+	-	-	-	-	-	-	-
<u>SMALL INTESTINE</u>	-	-	-	+	-	-	-	-	-
<u>PANCREAS</u>	-	-	-	-	-	-	-	-	-
<u>LARGE INTESTINE</u>	-	+	-	-	-	-	-	-	-
<u>Nematodiasis</u>									
<u>Enteritis, Acute</u>									
<u>LIVER</u>	-	-	-	-	-	-	-	-	-
<u>Hepatitis, Focal</u>	1								
<u>Infarct, Lobar</u>									
<u>ADRENAL</u>	-	-	-	-	-	-	-	-	-

STRAIN OF RAT: Colony
 EXPOSURE: 13 wks
 DOSE LEVEL: Control
 AGENT: White Phosphorus/Fe't

TABLE 7 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
 SMOKE II

	MALE			FEMALE		
	HIGH	MED	LOW	HIGH	MED	LOW
	78-833	78-870	78-959	78-834	78-872	78-960
	A	B	C	A	B	C
THYROID	-	-	-	-	-	-
Degeneration	3	3	2	1	1	1
THYMUS	+	-	-	-	-	-
Hemorrhage, Focal	-	-	-	-	-	-
KIDNEY	-	-	-	-	-	-
Hydronephrosis				1		1
Nephritis, Interstitial					1	1
Glomerulonephritis					1	1
Tubular Mineralization, Focal				1	1	1
Tubular Dilatation, Focal				1	1	1
Proteinuria				1		
BLADDER				-	-	-
Perivasculitis				+	+	+
OVARY				+	+	+
UTERUS				+	+	+
Hysteritis				-	-	-

STRAIN OF RAT: Colony
 EXPOSURE: 13 weeks
 DOSE LEVEL: Control
 AGENT: White Phosphorus/Felt

TABLE 8 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS

	MALE			FEMALE			
	HIGH	MED	LOW	HIGH	MED	LOW	
	78-833	78-870	78-959	78-834	78-872	78-960	
	A	B	C	A	B	C	
MAMMARY GLAND	+	+	-	+	+	-	
Adenoma							P
Mastitis							
TESTES							
PROSTATE	+	+	-	-	+	+	
Prostatitis				2	2		
MARROW							
Granulopoiesis, Accelerated				2			
SPLEEN				-			
Hemosiderosis				-			
BRAIN				-			
Encephalitis, Pyogranulomatous				-			
Necrosis, Focal				-			
EYE				-			
PITUITARY				-			
Adenoma				-			

STRAIN OF RAT: Colony
 EXPOSURE: 13 wks
 DOSE LEVEL: High (1000 mg/m³)
 AGENT: White Phosphorus/Felt

TABLE 9 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
 SMOKE II

	MALE				FEMALE					
	78-831	78-874	78-875		A	B	C	D	A	B
*Animals exposed with this group but held for 8 days for physiology studies prior to necropsy; not included in figures at left or in study.					*	*			*	*
NASAL TURBinate	3/3	+	-	-	A	B	C	D	A	B
Rhinitis, Focal										
LARYNX	2/2	-	+	-					-	-
Laryngitis		3	3						+	+
Submucosal Gland Duct Ectasia									+	+
TRACHEA	0/3	+	+	+					3	
Tracheitis										
Submucosal Gland Duct Ectasia										
LUNGS	4/4	-	-	-					-	-
Pneumonia, Interstitial				7					2	
Pneumonia, Granulomatous	1				2				2	
Pneumonia, Purulent									2	
Congestion									1	
Peribronchiolar Lymphoid	2	2	1	1	2	1			1	
Aggregates									3	
Bronchitis										

STRAIN OF RAT: Colony
 EXPOSURE: 13 wks
 DOSE LEVEL: High (1000 mg/m³)
 AGENT: White Phosphorus/Felt

TABLE 10 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
SMOKE II

	MALE				FEMALE			
	78-831	78-832	78-874	78-875	78-831	78-832	78-874	78-875
	A	B	C	D	A	B	C	D
HEART	4/4	-	-	-	-	-	-	-
Myocardial Fibrosis, Focal								
Myocarditis, Focal								
ESOPHAGUS	3/3	-	+	-	-	+	-	-
STOMACH	4/4	-	-	-	-	-	-	-
SMALL INTESTINE	3/4	-	-	+	-	-	-	-
PANCREAS	4/4	-	-	-	-	-	-	-
LARGE INTESTINE	3/1	-	+	-	-	+	-	+
Nematodiasis								
Enteritis, Acute								
LIVER								
Hepatitis, Focal								
Infarct, Lobar								
ADRENAL	3/4	-	-	-	-	+	-	+

*Animals exposed with this group but held for 8 days for physiology studies prior to necropsy; not included in figures at left or in study.

STRAIN OF RAT: Colony
 EXPOSURE: 13 wks
 DOSE LEVEL: High (1000 mg./m³)
 AGENT: White Phosphorus/Feit

TABLE 11 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
 SMOKE II

	MALE				FEMALE									
	78-831	78-874	78-874	78-832					78-875					
	A	B	C	D	A	B	C	D	A	B	C	D	A	B
THYROID	3/2	-	-	+	-	+	-	+	-	+	-	+	-	-
Degeneration														
THYMUS	4/3	-	-	-	-	-	-	-	-	-	-	-	-	-
Hemorrhage, Focal														3
KIDNEY	4/4	-	-	-	-	-	-	-	-	-	-	-	-	-
Hydronephrosis														
Nephritis, Interstitial														
Glomerulonephritis														
Tubular Mineralization, Focal														
Tubular Dilatation, Focal														
Proteinuria														
BLADDER	4/4	-	-	-	-	-	-	-	-	-	-	-	-	-
Perivasculitis														
OVARY	0/3													
UTERUS	0/4													
Heteritis														

STRAIN OF RAT: Colony
 EXPOSURE: 13 wks
 DOSE LEVEL: High (1000 mg/m³)
 AGENT: White Phosphorus/Feit

TABLE 72 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
SHOE II

	MALE				FEMALE			
	A	B	C	D	A	B	C	D
MAMMARY GLAND	0/0				*	*	*	*
Adenoma					+	+	+	+
Mastitis								-
TESTES	4/0	-	-	-			+	+
PROSTATE	4/0	-	-	-			+	+
Prostatitis							-	-
MARROW	4/3	-	-	-			-	-
Granulopoiesis, Accelerated								
SPLEEN	4/4	-	-	-			-	-
Hemosiderosis	1	1	1	1			2	3
BRAIN	4/4	-	-	-			-	-
Encephalitis, Pyogranulomatous								
Necrosis, Focal								1
EYE	4/4	-	-	-			-	+
PITUITARY	3/3	-	-	-			-	+
Adenoma							-	-

STRAIN OF RAT: Colony
 EXPOSURE: 13 wks
 DOSE LEVEL: Medium (500 mg/m³)
 AGENT: White Phosphorus/Ferit

TABLE 13 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
 SHOKE II

	MALE	FEMALE					
		A	B	C	D	E	F
NASAL TURBinate	6/6	-	-	-	+	+	-
Rhinitis, Focal	4/6	-	-	-	-	-	-
LARYNX	0/0	+	+	+	+	+	+
Laryngitis							
Submucosal Gland Duct Ectasia							
TRACHEA	5/6	-	+	-	-	-	-
Tracheitis		1	2	2	2	3	3
Submucosal Gland Duct Ectasia							
LUNGS	6/6	-	-	-	-	-	-
Pneumonia, Interstitial							
Pneumonia, Granulomatous							
Pneumonia, Purulent							
Congestion							
Peribronchiolar Lymphoid		1	1	2	1	1	1
Aggregates							
Bronchitis							

STRAIN OF RAT: Colony
EXPOSURE: 13 wks
DOSE LEVEL: Medium (500 mg/m³)
AGENT: White Phosphorus/Feit

TABLE 14 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS SMOKE II

STRAIN OF RAT: Colony
 EXPOSURE: 13 wks
 DOSE LEVEL: Medium (500 mg/m³)
 AGENT: White Phosphorus/Felt

TABLE 15 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS

	MALE						FEMALE					
	78-869			78-871			78-871			78-871		
	A	B	C	D	E	F	A	B	C	D	E	F
THYROID	6/6	-	+	-	-	-	-	-	-	-	-	-
Regeneration	5/6	-	2	2	1	4	2	3	2	3	3	3
THYROID	5/5	-	-	-	+	-	-	+	-	-	-	-
KIDNEY	6/6	-	-	-	-	-	-	-	-	-	-	-
Hydronephrosis												
Nephritis, Interstitial												
Glomerulonephritis												
Tubular Mineralization, Focal							1	2	2	2	2	2
Tubular Dilatation, Focal												
Proteinuria							1	1	1	1	1	1
BLADDER	5/2	-	-	+	-	-	-	+	-	+	+	+
Perivasculitis												
OVARY		0/5					+	-	-	-	-	-
UTERUS		0/5					-	-	+	-	-	-
Metriritis							1	1	1	1	1	1

STRAIN OF RAT: Colony
 EXPOSURE: 13 wks
 DOSE LEVEL: Medium (50) mg/m³
 AGENT: White Phosphorus/Felt

TABLE 16 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS

		MALE						FEMALE					
		A	B	C	D	E	F	A	B	C	D	E	F
MAMMARY GLAND	6/6	-	-	-	-	-	-	-	-	-	-	-	-
Adenoma	0/3	-	-	-	-	-	-	-	-	-	-	-	-
Mastitis	-	-	-	-	-	-	-	-	-	-	-	-	-
TESTES	6/0	-	-	-	-	-	-	-	-	-	-	-	-
PROSTATE	6/0	-	-	-	-	-	-	-	-	-	-	-	-
Prostatitis	-	-	-	-	-	-	-	-	-	-	-	-	-
MARROW	5/6	-	+	-	-	-	-	-	-	-	-	-	-
Granulopoiesis, Accelerated	-	-	-	-	-	-	-	-	-	-	-	-	-
SPLEEN	6/5	-	-	-	-	-	-	-	-	-	-	-	-
Hemosiderosis	-	-	-	-	-	-	-	-	-	-	-	-	-
BRAIN	6/6	-	-	-	-	-	-	-	-	-	-	-	-
Encephalitis, Pyogranulomatous	-	-	-	-	-	-	-	-	-	-	-	-	-
Necrosis, Focal	-	-	-	-	-	-	-	-	-	-	-	-	-
EYE	6/5	-	-	-	-	-	-	-	-	-	-	-	-
PITUITARY	6/6	-	-	-	-	-	-	-	-	-	-	-	-
Adenoma	-	-	-	-	-	-	-	-	-	-	-	-	-

STRAIN OF RAT: Colony
 EXPOSURE: 13 wks
 DOSE LEVEL: LCW (200 mg/m³)
 AGENT: White Phosphorus/Felt

TABLE 17: MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
 SMOKE II

	MALE						FEMALE					
	A	B	C	D	E	F	A	B	C	D	E	F
NASAL TURBINATE	6/6	-	-	-	-	-	-	-	-	-	-	-
Rhinitis, Focal												
LARYNX	1/4	+	-	+	+	+	+	-	+	-	-	-
Laryngitis									2			
Submucosal Gland Duct Ectasia									2	1	2	2
TRACHEA	5/4	-	+	-	-	-	-	-	-	+	-	+
Tracheitis												
Submucosal Gland Duct Ectasia												
LUNGS	6/6	-	-	-	-	-	-	-	-	-	-	-
Pneumonia, Interstitial									1	1	2	
Pneumonia, Granulomatous												
Pneumonia, Purulent												
Congestion												
Peribronchial Lymphoid	1	1	1	1	1	2	1	1	2	1	1	1
Aggregates												
Bronchitis												

STRAIN OF RAT: Colony
 EXPOSURE: 13 wks
 DOSE LEVEL: Low (200 mg/m³)
 AGENT: White Phosphorus/Fe t

TABLE 15 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
 SOURCE II

	MALE						FEMALE					
	A	B	C	D	E	F	A	B	C	D	E	F
HEART	6/6	-	-	-	-	-	-	-	-	-	-	-
Myocardial Fibrosis, Focal	-	-	-	-	-	-	-	-	-	-	-	-
Myocarditis, Focal	-	-	-	-	-	-	-	-	-	-	-	-
ESOPHAGUS	5/4	-	-	+	-	-	-	-	-	-	-	-
STOMACH	6/6	-	-	-	-	-	-	-	-	-	-	-
SMALL INTESTINE	6/6	-	-	-	-	-	-	-	-	-	-	-
PANCREAS	6/6	-	-	-	-	-	-	-	-	-	-	-
LARGE INTESTINE	6/6	-	-	-	-	-	-	-	-	-	-	-
Nematodiasis	-	-	-	-	-	-	-	-	-	-	-	-
Enteritis, Acute	-	-	-	-	-	-	-	-	-	-	-	-
LIVER	6/6	-	-	-	-	-	-	-	-	-	-	-
Hepatitis, Focal	-	-	-	-	-	-	-	-	-	-	-	-
Infarct, Lobar	-	-	-	-	-	-	-	-	-	-	-	-
ADRENAL	6/6	-	-	-	-	-	-	-	-	-	-	-

STRAIN OF RAT: Colony
 EXPOSURE: 13 V/IS
 DOSE LEVEL: Low (200 m3/m³)
 AGENT: White Phosphorus/Fer.t

TABLE 79 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS

	MALE						FEMALE					
	A	B	C	D	E	F	A	B	C	D	E	F
THYROID	4/4	-	+	-	-	-	-	+	-	-	-	+
Degeneration				2	2	2	2	1	1	2		
THYMUS	6/6	-	-	-	-	-	-	-	-	-	-	-
Hemorrhage, Focal												
KIDNEY	6/6	-	-	-	-	-	-	-	-	-	-	-
Hydronephrosis												
Nephritis, Interstitial												
Glomerulonephritis												
Tubular Mineralization, Focal												
Tubular Dilatation, Focal							1					
Proteinuria							1					
BLADDER	5/4	-	-	-	-	-	-	+	-	+	-	-
Perivasculitis												
OVARY	0/6											
UTERUS	0/5											
Menstritis												

STRAIN OF RAT: Colony
 EXPOSURE: 13 wks
 DOSE LEVEL: Low (200 mg/m³)
 AGENT: White Phosphorus/Feit

TABLE 20 : MICROSCOPIC OBSERVATIONS IN TISSUES FROM COLONY RATS
SMOKER 11

Appendix E

	MALE						FEMALE					
	A	B	C	D	E	F	A	B	C	D	E	F
MAMMARY GLAND	0/3											
Adenoma												
Mastitis												
TESTES	6/0											
PROSTATE	5/0	-	-	-	-	-						
Prostatitis												
MARROW	6/6	-	-	-	-	-						
Granulopoiesis, Accelerated						2						
SPLEEN	6/6	-	-	-	-	-						
Hemosiderosis												
BRAIN	6/6	-	-	-	-	-						
Encephalitis, Prgranulomatous												
Necrosis, Focal												
Eye	6/6	-	-	-	-	-						
PITUITARY												
Adenoma												

101 / 102

APPENDIX F

PATHOLOGY REPORT - 13-WEEK EXPOSURE AND 4-WEEK RECOVERY PERIOD

PATHOLOGY REPORT
PROJECT SMOKE II, WHITE PHOSPHORUS/FELT
INHALATION STUDY PERFORMED FOR THIRTEEN WEEKS IN
EDGEWOOD COLONY RATS
PROTOCOL NO. PEM 78-7

1. INTRODUCTION.

Male and female Edgewood area colony rats approximately six weeks old were exposed to high (1000 mg/m^3), medium (500 mg/m^3) and low (200 mg/m^3) dose levels of the agent White Phosphorus/Felt in the form of smoke. The above doses were administered 15 minutes a day, 5 days a week for a period of thirteen weeks. Thirty days following final exposure, the rats were killed by intraperitoneal injection of sodium pentobarbital and necropsied. Since each dose group was exposed on different calendar dates, separate control groups were utilized with each exposure group. Controls were placed in the exposure chamber for the same length of time each day as the exposed animals. Controls were necropsied concurrently with the exposed groups. All animals were housed in Building E3266 in standard laboratory animal facilities during non-exposure periods. Following necropsy, tissues were fixed in ten percent buffered formalin. The preserved tissues were submitted to the American HistoLab, for imbedding in paraffin, processing, and staining with hematoxylin and eosin. The following tissues from the high dose group and controls were processed for microscopic examination: nasal turbinates, larynx/trachea, lungs, heart, esophagus, salivary gland, stomach, small intestine, pancreas, large intestine, liver, adrenal, thyroid, thymus, kidney, bladder, ovary/testis, uterus, mammary gland, skin, muscle, prostate, bone marrow, bone, spleen, nerve, eye, brain, and pituitary. Turbinates were removed at necropsy for processing. The trachea was sampled from the proximal portion and lungs were sectioned randomly. The respiratory tracts only (target organ) consisting of nasal turbinates, trachea, larynx, and lungs were examined in medium and low dose groups.

2. RESULTS.

Significant findings were limited to the respiratory tract. These data are presented in Tables 1-3. Microscopic diagnoses from all tissues of high dose and control rats are presented in Tables 4-7. Respiratory tract lesions are presented from all other animals in Tables 8-13. Lesions were noted in the larynx or trachea of 15 of 16 high (1000 mg/m^3) dose level rats, 20 of 24 medium dose (500 mg/m^3) level rats and in none of the controls. Pulmonary lesions were noted in 11 of 16 high dose rats, 6 of 24 medium dose rats and none of the controls for these two groups. Although none of the low dose animals exhibited significant lesions, one control (79-0176) for this group did display the same pulmonary lesion as the high and medium dose exposed animals. Lesions were most extensive and severe in the larynx and trachea. They consisted mainly of thickening of the lamina propria and submucosa by collagen, endothelial cell proliferation, and macrophage infiltration. Areas of collagen were tinctorially different and characterized by an amphophilic to basophilic appearance. Epithelioid macrophages with giant cell formation surrounded some of the altered collagen. Overlying epithelium often lacked cilia and was at times thickened and metaplastic. Occasionally, inflammatory cells were seen

in the epithelium. Similar but much less extensive lesions were seen in bronchi and bronchioles. These lesions were characterized by focal areas of altered collagen within the lamina propria. These foci were amphophilic to basophilic in appearance and occasionally surrounded by macrophages and epithelioid giant cells. Small granulomas were noted within respiratory bronchioles. These granulomas often extended into the adjacent lung parenchyma. The granulomas were characterized by swirling configurations of histiocytes and epithelioid macrophages with giant cell formation.

3. DISCUSSION.

A number of incidental lesions were observed in various organs of the high (1000 mg/m^3) dose group. These lesions either occurred sporadically or with equal frequency in both exposed and control animals. Smoke-related lesions were limited to the upper airways and to a lesser extent, the smaller airways of the lung and the parenchyma adjacent to the terminal bronchioles. Since only the respiratory system was involved in the high dose animals, examination was confined to this system in the medium and low dose groups. All lesions attributable to the smoke were of a chronic inflammatory nature, characterized by collagen degeneration, and thickening of the lamina propria and submucosa of airways by fibrovascular proliferations.

Nasal lesions consisting of focal squamous metaplasia and a slight inflammatory cell infiltrate were seen in one high dose animal, accession number 78-910F. It is felt that lesions may have been present in other exposed animals as well; however, the sampling technique precluded our observing this. In light of this, it is recommended that a more systematic approach to studying the upper respiratory tract lesions be followed in subsequent studies.

4. CONCLUSION.

Rats exposed to the agent, White Phosphorus/Felt at the 1000 mg/m^3 and 500 mg/m^3 dosage levels developed a dosage related response characterized by chronic inflammation of the airways; lesions were most severe in the larynx and trachea but also involved the lower areas to a slight degree.

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30 May 1980

REVIEWED BY:

W. Hall
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U.S. Army
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TABLE 1 MICROSCOPIC OBSERVATIONS FROM RESPIRATORY TISSUES FROM
EDGEWOOD AREA COLONY RATS EXPOSED TO WHITE PHOSPHORUS/FELT
SMOKE FOR 13 WEEKS AND KILLED 30 DAYS POST EXPOSURE

PROTOCOL NO. PEM 78-7	SEX	MALE	FEMALE
	DOSE	1000PPM	CONTROL
	ACCESSION NO.	78-910	78-912
NO. OF RATS	9	6	7
DIAGNOSIS			
Laryngitis/Tracheitis	9	0	6
Bronchitis/Bronchiolitis	4	0	2
Granulomatous Bronchiolitis/Pneumonia	3	0	2

PROTOCOL NO. PEM 78-7

TABLE 2 MICROSCOPIC OBSERVATIONS FROM RESPIRATORY TISSUES FROM
EDGWOOD AREA COLONY RATS EXPOSED TO WHITE PHOSPHORUS/FELT
SMOKE FOR 13 WEEKS AND KILLED 30 DAYS POST EXPOSURE

SEX	MALE			FEMALE		
	DOSE	500PPM	CONTROL	500PPM	CONTROL	
ACCESSION NO.		78-926	78-927	78-929	78-930	
NO. OF RATS		12	6	12	6	
<u>DIAGNOSIS</u>						
Laryngitis/Tracheitis	10	0	10	0	0	
Bronchitis/Bronchiolitis	4	0	1	1	0	
Granulomatous	0	0	1	1	0	
Bronchitis/Pneumonia						

PROTOCOL NO. PEM 78-7

TABLE 3 MICROSCOPIC OBSERVATIONS FROM RESPIRATORY TISSUES FROM
EDGEWOOD AREA COLONY RATS EXPOSED TO WHITE PHOSPHORUS/FELT
SMOKE FOR 13 WEEKS AND KILLED 30 DAYS POST EXPOSURE

SEX	MALE		FEMALE	
	DOSE	200PPM	DOSE	200PPM
ACCESSION NO.	79-017		79-016	79-018
NO. OF RATS	6	3	6	2
<u>DIAGNOSIS</u>				
Laryngitis/Tracheitis	0	0	0	0
Bronchitis/Bronchiolitis	0	0	0	0
Granulomatous Bronchiolitis/Pneumonia	0	0	0	0

PROTOCOL NO.: PEM 78-7
SEX: MALE
EXPOSURE: CONTROL
AGENT: WHITE PHOSPHORUS/FELT
LENGTH OF EXPOSURE: 13 WEEKS
POST EXPOSURE: 30 DAYS

Appendix F

TABLE 4 MICROSCOPIC OBSERVATIONS IN TISSUES FROM EDGEWOOD AREA COLONY RATS
SMOKE II

<u>ACCESSION NO.</u>	<u>DIAGNOSIS</u>
78-912A	Histiocytosis, focal, minimal, pleura, lung, rat Heprosis, multifocal, minimal to mild, kidney Congestion, minimal, thymus
78-912B	Congestion, mild, lung, rat Hemorrhage and congestion, minimal, thymus
78-912C	Pyelitis, focal, acute, minimal, kidney, rat Hemorrhage and congestion, diffuse, mild, thymus
78-912D	Congestion, multifocal, mild, lung, rat Chronic respiratory disease, minimal, lung & bronchus Congestion and hemorrhage, diffuse, minimal, thymus
78-912E	Congestion, multifocal, mild, lung, rat Chronic respiratory disease, minimal, lung Hemorrhage, multifocal, minimal, thymus
78-912F	Lymphoid infiltrate focal, minimal, subplura, lung, rat Chronic respiratory disease, minimal, lung Congestion, diffuse, minimal, thymus Hemorrhage, multifocal, minimal, thymus Medial calcification, focal, minimal, pulmonary artery

PROTOCOL NO.: PEM 78-7
SEX: FEMALE
EXPOSURE: CONTROL
AGENT: WHITE PHOSPHORUS/FELT
LENGTH OF EXPOSURE: 13 WEEKS
POST EXPOSURE: 30 DAYS

Appendix F

TABLE 5 MICROSCOPIC OBSERVATIONS IN TISSUES FROM EDGEWOOD AREA COLONY RATS
SMOKE II

<u>ACCESSION NO.</u>	<u>DIAGNOSIS</u>
78-914A	Histiocytosis, focal, minimal, lung, rat Nephrocalcinosis, minimal, kidney Hemorrhage, multifocal, minimal, thymus
78-914B	Congestion, multifocal, mild, lung, rat Nephrocalcinosis, minimal, kidney Hemorrhage, multifocal, minimal, thymus
78-914C	Nephrocalcinosis, minimal, kidney, rat
78-914D	Congestion, multifocal, mild, lung, rat Retinal atrophy, focal, minimal, eye Endocardiosis, valvular, minimal, heart
78-914E	Tracheitis, focal, lymphocytic, minimal, trachea, rat Chronic respiratory disease, minimal, lung Congestion, focal, minimal, lung Granuloma, focal, Hardarian gland Congestion, diffuse, moderate, thymus

PROTOCOL NO.: PEM 78-7
SEX: FEMALE
EXPOSURE: CONTROL
AGENT: WHITE PHOSPHORUS/FELT
LENGTH OF EXPOSURE: 13 WEEKS
POST EXPOSURE: 30 DAYS

TABLE 5 MICROSCOPIC OBSERVATIONS IN TISSUES FROM EDGEWOOD AREA COLONY RATS
SMOKE II

Appendix F

ACCESSION NO.

78-914F

DIAGNOSIS

Congestion, focal, minimal, lung, rat

Congestion and hemorrhage, diffuse, mild, thymus

Congestion, moderate, lymph node

PROTOCOL NO.: PEM 78-7
 SEX: MALE
 EXPOSURE: 1000 MG/M³
 AGENT: WHITE PHOSPHORUS/FELT
 LENGTH OF EXPOSURE: 13 WEEKS
 POST EXPOSURE: 30 DAYS

Appendix F

TABLE 6 MICROSCOPIC OBSERVATIONS IN TISSUES FROM EDGEWOOD AREA COLONY RATS
SMOKE II

ACCESSION NO.

78-910A

DIAGNOSIS

Tracheitis, chronic, diffuse, moderate, trachea, rat
 Interstitial pneumonia, multifocal, mild, lung
 Congestion, diffuse, minimal, spleen

 Tracheitis, chronic, diffuse, mild, trachea, rat
 Prostatitis, interstitial, subacute, diffuse, minimal-mild, prostate
 Epicarditis, subacute, focal, mild, rt. ventricle, heart

 Tracheitis, chronic, diffuse, mild, trachea, rat
 Prostatitis, interstitial, subacute, diffuse, minimal, prostate
 Nematodiasis, large intestine, compatible with Oxyuriasis

 Tracheitis, subacute, diffuse, moderate-severe, trachea, rat
 Pneumonia, granulomatous, multifocal, minimal, lung
 Prostatitis, subacute, multifocal, minimal, prostate
 Congestion, diffuse, minimal, thymus

PROTOCOL NO.: PEM 78-7
SEX: MALE
EXPOSURE: 1000 MG/M³
AGENT: WHITE PHOSPHORUS/FELT
LENGTH OF EXPOSURE: 13 WEEKS
POST EXPOSURE: 30 DAYS

Appendix F

TABLE 6 MICROSCOPIC OBSERVATIONS IN TISSUES FROM EDGEWOOD AREA COLONY RATS
SMOKE 1!

ACCESSION NO.

78-910E

DIAGNOSIS

Tracheitis, chronic, multifocal, mild, trachea, rat
Laryngitis, chronic, multifocal, mild, larynx
Bronchitis, chronic, multifocal, mild, lung
Rhinitis, focal, subacute, mild, nasal cavity
Myocarditis, focal, subacute, minimal, heart
Congestion, diffuse, mild, spleen

78-910F

Tracheitis, multifocal, chronic, mild, trachea, rat
Pneumonia, granulomatous, multifocal, minimal, lung
Bronchitis, multifocal, chronic, minimal, lung
Prostatitis, interstitial, multifocal, minimal, prostate
Congestion, and hemorrhage, diffuse, mild, thymus

78-910G

Tracheitis, multifocal, chronic, minimal, trachea, rat
Bronchiolitis, chronic, multifocal, minimal, lung
Prostatitis, subacute, multifocal, mild, prostate

78-910H

Laryngitis, chronic, diffuse, minimal, larynx, rat
Pneumonia, granulomatous, multifocal, minimal, lung
Hemorrhage, focal, minimal, thymus, rat

PROTOCOL NO.: PEM 78-7
SEX: MALE
EXPOSURE: 1000 MG/M³
AGENT: WHITE PHOSPHORUS/FELT
LENGTH OF EXPOSURE: 13 WEEKS
POST EXPOSURE: 30 DAYS

TABLE 6 MICROSCOPIC OBSERVATIONS IN TISSUES FROM EDGEWOOD FRESH CORONY RATS
SMOKE !!

ACCESSION NO.

78-9101

DIAGNOSIS

Laryngitis, diffuse, chronic, mild, larynx, rat
Pneumonia, granulomatous, multifocal, minimal, lung
Bronchiolitis, obliterative, focal, lung
Bronchiolitis, multifocal, chronic, minimal
Congestion, diffuse, mild, thymus
Congestion, diffuse, minimal, spleen

PROTOCOL NO.: PEM 78-7
 SEX: FEMALE
 EXPOSURE: 1000 MG/M³
 AGENT: WHITE PHOSPHORUS/FELT
 LENGTH OF EXPOSURE: 13 WEEKS
 POST EXPOSURE: 30 DAYS

TABLE 7 MICROSCOPIC OBSERVATIONS IN TISSUES FROM EDGEWOOD AREA COLONY RATS
SMOKE II

<u>ACCESSION NO.</u>	<u>DIAGNOSIS</u>
78-913A	Tracheitis, chronic, multifocal, mild, trachea, rat Hemorrhage, multifocal, minimal, thymus Congestion, multifocal, mild, lung Nephrocalcinosis and lithiasis, mild, kidney
78-913B	Laryngitis, multifocal, chronic, mild, larynx, rat Bronchitis, multifocal, chronic, minimal, lung Pneumonia, granulomatous, focal, minimal, lung Nephrocalcinosis, minimal, kidney Atrophy, focal, minimal, retina, eye Congestion, and hemorrhage, diffuse, minimal, thymus
78-913C	Bronchitis, focal, chronic, minimal, lung, rat Congestion, and hemorrhage, diffuse, mild, thymus Myocarditis, focal, subacute, minimal, Rt. ventricle, heart Nephrocalcinosis, minimal, kidney

PROTOCOL NO.: PEM 78-7
SEX: FEMALE
EXPOSURE: 1000 MG/M³
AGENT: WHITE PHOSPHORUS/FELT
LENGTH OF EXPOSURE: 13 WEEKS
POST EXPOSURE: 30 DAYS

TABLE 7 MICROSCOPIC OBSERVATIONS IN TISSUES FROM EDGEWOOD AREA COLONY RATS
SMOKE II

ACCESSION NO.

78-913D

DIAGNOSIS

Laryngitis, subacute, multifocal, minimal, larynx, rat
Histiocytosis, multifocal, minimal, lung
Hemorrhage, diffuse, moderate, lymph node
Congestion, and hemorrhage, diffuse, mild, thymus

Tracheitis, chronic, focal, minimal, trachea, rat
Lymphoid infiltrate: focal, minimal, subpleural, lung
Histiocytosis, focal, minimal, lung, rat
Congestion, and hemorrhage, diffuse, mild, thymus
Nephrocalcinosis, minimal, kidney
Mineralization, focal, minimal, adrenal

78-913E
78-913F

Tracheitis, focal, chronic, minimal, trachea, rat
Congestion, diffuse, mild, thymus
Cyst, thymus
Hemorrhage, diffuse, moderate, lymph node
Nephrocalcinosis, mild, kidney

PROTOCOL NO.: PEM 78-7
SEX: FEMALE
EXPOSURE: 1000 MG/M³
AGENT: WHITE PHOSPHORUS/FELT
LENGTH OF EXPOSURE: 13 WEEKS
POST EXPOSURE: 30 DAYS

Appendix F

TABLE 7 MICROSCOPIC OBSERVATIONS IN TISSUES FROM EDGEWOOD AREA COLONY RATS SMOKE II

ACCESSION NO.

78-9136

DIAGNOSIS

Laryngitis, multifocal, chronic, minimal, larynx, rat
Pneumonia, granulomatous, multifocal, minimal, lung
Chronic respiratory disease, minimal
Congestion, diffuse, mild, lung
Medial calcification, multifocal, minimal, pulmonary artery
Hemorrhage and congestion, thymus

PROTOCOL NO.: PEM 78-7
SEX: MALE
EXPOSURE: CONTROL
AGENT: WHITE PHOSPHORUS/FELT
LENGTH OF EXPOSURE: 13 WEEKS
POST EXPOSURE: 30 DAYS

TABLE 8 MICROSCOPIC OBSERVATIONS IN RESPIRATORY TISSUES FROM EDGEWOOD AREA COLONY RATS
SMOKE II

<u>ACCESSION NO.</u>	<u>DIAGNOSIS</u>
78-927A	Congestion, mild, lung, rat
78-927B	Congestion, mild, lung, rat
78-927C	No significant lesion
78-927D	Medial calcification, focal, minimal, pulmonary artery, rat
78-927E	Congestion, mild, lung, rat Medial calcification, multifocal, minimal, pulmonary artery Chronic respiratory disease, mild, lung
78-927F	Lymphoid infiltrate, focal, minimal, pleura, lung, rat Congestion, minimal, lung

PROTOCOL NO.: PEM 78-7
SEX: MALE
EXPOSURE: 500 MG/M³
AGENT: WHITE PHOSPHORUS/FELT
LENGTH OF EXPOSURE: 13 WEEKS
POST EXPOSURE: 30 DAYS

TABLE 9 MICROSCOPIC OBSERVATIONS IN RESPIRATORY TISSUES FROM EDGEWOOD AREA COLONY RATS
SMOKE II

ACCESSION NO.

78-926A

Tracheitis, chronic, diffuse, mild, trachea, rat
Bronchiolitis, focal, chronic, minimal, lung
Medial calcification, focal, minimal, pulmonary artery, lung

78-926B

No significant lesion

78-926C

Tracheitis, diffuse, chronic, moderate, trachea, rat
Bronchitis, focal, chronic, minimal, lung

78-926D

Tracheitis, diffuse, chronic, moderate, trachea, rat
Bronchitis, multifocal, chronic, minimal, bronchi
Chronic respiratory disease, minimal, lung
Medial calcification, multifocal, minimal, pulmonary artery, lung

78-926E

Tracheitis, focal, chronic, moderate, trachea, rat
Medial hypertrophy, multifocal, moderate, pulmonary artery, lung

78-926F

Laryngitis, diffuse, chronic, mild, larynx, rat

PROTOCOL NO.: PEM 78-7 TABLE 9 MICROSCOPIC OBSERVATIONS IN RESPIRATORY TISSUES FROM EDGEWOOD AREA COLONY RATS

SEX: MALE

EXPOSURE: 500 MG/M³

AGENT: WHITE PHOSPHORUS/FELT

LENGTH OF EXPOSURE: 13 WEEKS

POST EXPOSURE: 30 DAYS

Appendix F

<u>ACCESSION NO.</u>	<u>DIAGNOSIS</u>
78-926G	Tracheitis, diffuse, chronic, moderate, trachea, rat
78-926H	Bronchiolitis, focal, chronic, minimal, lung, rat
78-926I	Tracheitis, multifocal, chronic, moderate, trachea, rat Histiocytosis, multifocal, minimal, lung
78-926J	Laryngitis, multifocal, chronic, minimal, larynx, rat Medial calcification, focal, minimal, pulmonary artery, lung
78-926K	Tracheitis, multifocal, chronic, moderate, trachea, rat
78-926L	Tracheitis, focal, chronic, mild, trachea, rat

PROTOCOL NO.: PEM 78-7 TABLE 10 MICROSCOPIC OBSERVATIONS IN RESPIRATORY TISSUES FROM EDGENWOOD AREA COLONY RATS
SEX: FEMALE EXPOSURE: 500 MG/M³

AGENT: WHITE PHOSPHORUS/FELT
LENGTH OF EXPOSURE: 13 WEEKS
POST EXPOSURE: 30 DAYS

Appendix F

<u>ACCESSION NO.</u>	<u>DIAGNOSIS</u>
78-929A	Tracheitis, multifocal, chronic, minimal, trachea, rat Bronchiolitis, granulomatous, focal, minimal, lung
78-929B	Interstitial pneumonia, focal, minimal, lung, rat
78-929C	Essentially normal tissues, rat
78-929D	Tracheitis, focal, chronic, minimal, trachea, rat
78-929E	Laryngitis, focal, chronic, minimal, larynx, rat
78-929F	Tracheitis, diffuse, chronic, moderate, trachea, rat
78-929G	Laryngitis, focal, chronic, minimal, larynx, rat
78-929H	Laryngitis, focal, chronic, minimal, trachea, rat
78-929I	Laryngitis, focal, chronic, mild, larynx, rat
78-929J	Tracheitis, multifocal, chronic, minimal, trachea, rat

PROTOCOL NO.: PEM 78-7 TABLE 10 MICROSCOPIC OBSERVATIONS IN RESPIRATORY TISSUES FROM EDGEWOOD AREA COLONY RATS

SEX: FEMALE
EXPOSURE: 500 MG/M³
AGENT: WHITE PHOSPHORUS/FELT
LENGTH OF EXPOSURE: 13 WEEKS
POST EXPOSURE: 30 DAYS

Appendix F

SMOKE 11

ACCESSION NO.

78-929K

78-929L

DIAGNOSIS

Tracheitis, multifocal, chronic, minimal, trachea, rat
Laryngitis, diffuse, chronic, minimal, larynx, rat
Bronchiolitis, focal, chronic, minimal, lung

PROTOCOL NO.: PEM 78-7 TABLE 11 MICROSCOPIC OBSERVATIONS IN RESPIRATORY TISSUES FROM EDGEWOOD AREA COLONY RATS

SEX: FEMALE

EXPOSURE: CONTROL

AGENT: WHITE PHOSPHORUS/FELT

LENGTH OF EXPOSURE: 13 WEEKS

POST EXPOSURE: 30 DAYS

Appendix F

ACCESSION NO.

78-930A

DIAGNOSIS

Histiocytosis, focal, minimal, lung, rat
Chronic respiratory disease, minimal, lung

Chronic respiratory disease, minimal, lung, rat

78-930B

Histiocytosis, multifocal, mild, lung, rat
Chronic respiratory disease, lung

78-930D

Histiocytosis, multifocal, minimal, lung, rat
Chronic respiratory disease, minimal, lung

78-930E

Histiocytosis, multifocal, minimal, lung, rat
Chronic respiratory disease, lung

PROTOCOL NO.: PEM 78-7 TABLE 12 MICROSCOPIC OBSERVATIONS IN RESPIRATORY TISSUES FROM EDGEWOOD AREA COLONY RATS
SEX: MALE 200 MG/M³
EXPOSURE:
AGENT: WHITE PHOSPHORUS/FELT
LENGTH OF EXPOSURE: 13 WEEKS
POST EXPOSURE: 30 DAYS

TABLE 12 MICROSCOPIC OBSERVATIONS IN RESPIRATORY TISSUES FROM EDGEWOOD AREA COLONY RATS

Appendix F

<u>ACCESSION NO.</u>	<u>DIAGNOSIS</u>
79-15A	No significant lesions
79-15B	No significant lesions
79-15C	No significant lesions
79-15D	No significant lesions
79-15E	No significant lesions
79-15F	No significant lesions

PROTOCOL NO.: PEM 78-7 TABLE 13 MICROSCOPIC OBSERVATIONS IN RESPIRATORY TISSUES FROM EDGEWOOD AREA COLONY RATS
SEX: MALE
EXPOSURE: CONTROL
AGENT: WHITE PHOSPHORUS/FELT
LENGTH OF EXPOSURE: 13 WEEKS
POST EXPOSURE: 30 DAYS

Appendix F

SMOKE II

ACCESSION NO.

79-17A

Histiocytosis, focal, minimal, lung, rat
Congestion, minimal, lung
Chronic respiratory disease, minimal, lung
Medial calcification, focal, minimal, pulmonary artery, lung

79-17B

Chronic respiratory disease, minimal, lung, rat
Congestion, minimal, lung

79-17C

Pneumonia, granulomatous, multifocal, minimal, lung, rat
Chronic respiratory disease, lung

DIAGNOSIS

Histiocytosis, focal, minimal, lung, rat

Congestion, minimal, lung

Chronic respiratory disease, minimal, lung

Medial calcification, focal, minimal, pulmonary artery, lung

Chronic respiratory disease, minimal, lung, rat

Congestion, minimal, lung

Pneumonia, granulomatous, multifocal, minimal, lung, rat

Chronic respiratory disease, lung

PROTOCOL NO.: PEM 78-7 TABLE 14 MICROSCOPIC OBSERVATIONS IN RESPIRATORY TISSUES FROM EDGEWOOD AREA COLONY RATS
 SEX: FEMALE
 EXPOSURE: 200 MG/M³
 AGENT: WHITE PHOSPHORUS/FELT
 LENGTH OF EXPOSURE: 13 WEEKS
 POST EXPOSURE: 30 DAYS

Appendix F

<u>ACCESSION NO.</u>	<u>DIAGNOSIS</u>
79-16A	Histiocytosis, focal, minimal, lung, rat Pneumonia, interstitial, focal, minimal, lung
79-16B	Congestion, minimal, lung, rat
79-16C	Histiocytosis, multifocal, minimal, lung, rat Lymphoid infiltrate, focal, minimal, lung
79-16D	Congestion, mild, lung, rat
79-16E	Medial calcification, focal, minimal, pulmonary artery, lung, rat
79-16F	Histiocytosis, multifocal, minimal, lung, rat

PROTOCOL NO.: PFM 78-7 TABLE 15 :MICROSCOPIC OBSERVATIONS I.4 RESPIRATORY TISSUES FROM EDGEWOOD AREA COLOR RATS

SEX: FEMALE

EXPOSURE: CONTROL

AGENT: WHITE PHOSPHORUS/FELT

LENGTH OF EXPOSURE: 13 WEEKS

POST EXPOSURE: 30 DAYS

Appendix F

ACCESSION NO.

79-18A

79-16B

DIAGNOSIS

Chronic respiratory disease, minimal, lung, rat

Congestion, minimal, lung, rat

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